REFERENCE: Derwent DGene Search Report

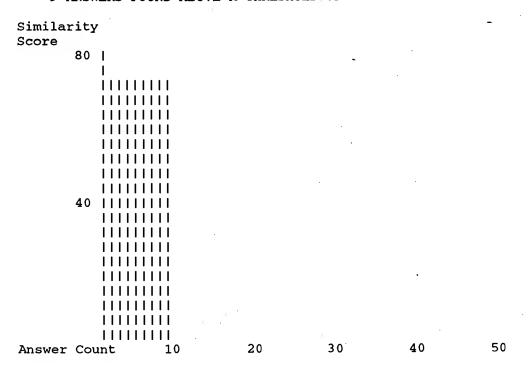
FRANZ-BACON, et al., USSN: 09/099,898

Atty. Docket No.: DX0744K

Mouse C2

 ${\tt MKTTTCSLLICISLLQLMVPVNTDETIEIIVENKVKELLANPANYPSTVTKTLSCTSVKTMNRWASCPAGMTATGCACGFACGSWEIQSGDTCNCLCLLVDWTTARCCQLS}$

9 ANSWERS FOUND ABOVE A THRESHOLD OF 67



L2 ANSWER 1 OF 9 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 96P-R98208 Protein DGENE

TITLE: Cell-targetted retroviral vector particles - having

envelope protein modified with targetting polypeptide

INVENTOR: Anderson W; Chiang Y L; Januszeski M; Mackrell A J;

Zhao Y

PATENT ASSIGNEE: (GENE-N)GENETIC THERAPY INC

(UYSC-N) UNIV SOUTHERN CALIFORNIA

PATENT INFO: WO 9630504 A1 961003 - 73 pp

APPLICATION INFO: WO 96-US3908 960322 PRIORITY INFO: US 95-409648 950324 PAT. SEQ. LOC: Example 2; Page 36

DATA ENTRY DATE: 30 DEC 1996 (first entry)

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 96-455352 [45]

DESCRIPTION: Nucleotide used in production of MSH/MoMuLV chimeric

sequence

KEYWORD: Moloney murine leukaemia virus; gp70; 4070A retrovirus;

retrovirus; 10A1 murine leukaemia virus; NZB-9-1 murine leukaemia virus; polytropic MX27 provirus; targetted drug delivery; gene therapy; single chain antibody;

envelope protein; ss

ORGANISM: Synthetic

ABSTRACT:

Cell targetted retroviral vector particles can be used in gene therapy to deliver a heterologous gene to a target cell for expression of a heterologous polypeptide in that cell. The cell targetted retroviral vector particles comprise an envelope protein which is modified to contain a targetting polypeptide (a single chain antibody), or in the case of moloney murine leukaemia virus (MoMuLV), alpha melanotropin-stimulating hormone (MSH). Two oligonucleotides (R98207, R98208) were used to substitute sequences in MoMuLV for MSH sequences. This oligonucleotide was used to replace residues G80-P88 of MoMuLV envelope protein (See W04248)

AMINO ACID COUNTS:8 A; 0 R; 0 N; 0 D; 0 B; 17 C; 0 Q; 0 E; 0 Z; 8 G; 0 H; 0 I; 0 L; 0 K; 0 M; 0 F; 0 P; 0 S; 11

T; 0 W; 0 Y; 0 V;

SEQUENCE LENGTH: 44

SEQUENCE

1 catttccgat ggtgcaagcc ggtattaacc tccctcaccc ctcg

ALIGN Smith-Waterman score: 86
43 aa overlap starting at 5

scpagmtatgcacgfacgsweiqsgdtcnclcllvdwttarcc

tccgatggtgcaag_ccgg_tattaacctccc____tcacc

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

```
ANSWER 2 OF 9 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 98P-W56120 protein
                  Reducing severity of host versus graft reaction -
TITLE:
                  comprises suppressing auto-immunity to heat-shock
                  protein to prevent rejection
                  Birk O; Cohen I R
INVENTOR:
                  (YEDA) YEDA RES & DEV CO LTD
PATENT ASSIGNEE:
                  WO 9808536 A2 980305
                                                     37 pp
PATENT INFO:
APPLICATION INFO: WO 97-US15294 970902
                  US 96-706209
                                 960830
PRIORITY INFO:
                  Disclosure; Pages 28-29
PAT. SEQ. LOC:
DATA ENTRY DATE: 08 JUL 1998 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
                  98-179175 [16]
OTHER SOURCE:
                  Protein sequence of human heat shock protein (hsp) 60
DESCRIPTION:
                  Heat shock protein; hsp60; human; host versus graft
KEYWORD:
                  reaction; HVGR; transplantation; organ; tissue;
                  downregulatation; autoimmunity; prevention; graft
                  rejection
ORGANISM:
                  Homo sapiens
ABSTRACT:
      The present sequence represents human heat shock protein 60
      (hsp60). This protein is a stress protein expressible in all cells
      of the body. The severity of a host versus graft reaction (HVGR)
      concominant with transplantation of donor organ or tissue, can be
      reduced by downregulating hsp60 autoimmunity in the host. The
      specification also describes a method for selecting peptides for
      preventing or suppressing graft rejection. This method comprises
      treating a panel of labelled peptides with antigen-presenting cells
      isolated from peripheral blood lymphocytes of the candidate host,
      and selecting those that bind with the antigen presenting cell.
      hsp60, or its peptides, analogues, salts and functional derivatives
      can be used for downregulating hsp60 autoimmunity especially for
      reducing HVGR. hsp60 autoimmunity can accelerate foreign immunity
      and its downregulation helps prevent graft rejection
AMINO ACID COUNTS:57 A; 22 R; 20 N; 37 D; 0 B; 3 C; 17 Q; 43 E; 0 Z;
                  57 G; 3 H; 43 I; 48 L; 53 K; 18 M; 10 F; 19 P; 24 S;
                  34 T; 1 W; 7 Y; 57 V;
                  573
SEQUENCE LENGTH:
SEQUENCE
        1 mlrlptvfrq mrpvsrvlap hltrayakdv kfgadaralm lqgvdllada
       51 vavtmgpkgr tviieggwgs pkvtkdgvtv aksidlkdky knigaklvqd
      101 vanntneeag dgtttatvla rsiakegfek iskganpvei rrgvmlavda
      151 viaelkkqsk pyttpeeiaq vatisangdk eigniisdam kkygrkgyit
      201 vkdgktlnde leiiegmkfd rgyispyfin tskggkcefg dayvllsekk
      251 issiqsivpa leianahrkp lviiaedvdg ealstlvlnr lkvglqvvav
      301 kapgfgdnrk nqlkdmaiat ggavfgeegl tlnledvqph dlgkvgeviv
      351 tkddamllkg kgdkaqiekr iqeiieqldv ttseyekekl nerlaklsdg
      401 vavlkvggts dvevnekkdr vtdalnatra aveegivlgg gcallrcipa
      451 ldsltpaned gkigieiikr tlkipamtia knagvegsli vekimqssse
      501 vgydamagdf vnmvekgiid ptkvvrtall daagvasllt taevvvteip
      551 keekdpamaa maamaaamaa amf
ALIGN Smith-Waterman score: 81
      38 aa overlap starting at .442
      csllicisllqlmvpvntdetieiivenkvkellanpa
       :::: :: :: :::::: :: :: :: ::: ::
```

SEARCHED ON 26 OCT 1998

callrcipaldsltpanedq_kigiei_ikrtlkipa

```
ANSWER 3 OF 9 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 97P-W12345 protein
                                            DGENE
                 New peptide(s) derived from human heat-shock protein 60
TITLE:

    used for early diagnosis, prevention and treatment of

                  insulin-dependent diabetes mellitus
                  Abulafia R; Bockova J; Cohen I R; Elias D
INVENTOR:
                  (YEDA) YEDA RES & DEV CO LTD
PATENT ASSIGNEE:
                 WO 9701959 A1 970123
PATENT INFO:
                                                     49 pp
APPLICATION INFO: WO 96-US11375 960701
                  IL 95-114407
                                 950630
PRIORITY INFO:
                 Disclosure; Page 28-29
PAT. SEQ. LOC:
DATA ENTRY DATE: 13 NOV 1997 (first entry)
DOCUMENT TYPE:
                  Patent
                  English
LANGUAGE:
                  97-108693 [10]
OTHER SOURCE:
DESCRIPTION:
                 Human heat-shock protein 60
                  Heat-shock protein; hsp; hsp60; insulin-dependent
KEYWORD:
                  diabetes mellitus; IDDM
                  Homo sapiens
ORGANISM:
ABSTRACT:
      The peptides given in W12346 to W12358 are derived from human hsp60
      (W12345) and are useful for early diagnosis of IDDM by detecting,
      in the blood or urine, antibodies or T-cells immunologically
      reactive with human hsp60 (presence of which indicates high
      probability of diabetes or its subsequent development). Other
      peptides (W12359 to W12361) were shown not to be as effective
AMINO ACID COUNTS:57 A; 22 R; 20 N; 37 D; 0 B; 3 C; 17 Q; 43 E; 0
                  57 G; 3 H; 43 I; 48 L; 53 K; 18 M; 10 F; 19 P; 24 S;
                  34 T; 1 W; 7 Y; 57 V;
                  573
SEQUENCE LENGTH:
SEQUENCE
        l mlrlptvfrq mrpvsrvlap h1trayakdv kfgadaralm lqgvd11ada
       51 vavtmgpkgr tviieqgwgs pkvtkdgvtv aksidlkdky knigaklvqd
      101 vanntneeag dgtttatvla rsiakegfek iskganpvei rrgvmlavda
      151 viaelkkqsk pvttpeeiaq vatisangdk eigniisdam kkvgrkgvit
      201 vkdgktlnde leiiegmkfd rgyispyfin tskgqkcefq dayv11sekk
      251 issiqsivpa 1eianahrkp 1viiaedvdg ea1st1v1nr 1kvg1qvvav
      301 kapgfgdnrk nq1kdmaiat ggavfgeegl t1n1edvqph d1gkvgeviv
      351 tkddam11kg kgdkaqiekr iqeiieq1dv ttseyekek1 ner1ak1sdg
      401 vav1kvggts dvevnekkdr vtdalnatra aveegiv1gg gcallrcipa
      451 ldsltpaned qkigieiikr t1kipamtia knagvegsli vekimqssse
      501 vgydamagdf vnmvekgiid ptkvvrta11 daagvas11t taevvvteip
      551 keekdpgmga mggmgggmgg gmf
ALIGN Smith-Waterman score: 81
      38 aa overlap starting at 442
      cs11icis11qlmvpvntdetieiivenkvke11anpa
      :.:: :: :. ..:.: :. .: .: .: ::
```

SEARCHED ON 26 OCT 1998 FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

callrcipaldsltpanedq_kigiei_ikrt1kipa

```
ANSWER 4 OF 9 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 97P-W01657 Protein
                                            DGENE
                  Compsns. for treating or preventing insulin-dependent
TITLE:
                  diabetes - based on T cells specific for 65 kD
                  heat-shock protein
INVENTOR:
                  Cohen I R; Elias D; Markovits D
PATENT ASSIGNEE:
                  (YEDA) YEDA RES & DEV CO LTD
PATENT INFO:
                                                     29 pp
                  US 5578303 A 961126
APPLICATION INFO: US 89-322864
                                 890314
PRIORITY INFO:
                  US 91-751448
                                 910829
                  US 89-322864
                                 890314
                  US 89-371249
                                 890626
                  US 90-493127
                                 900314
                  US 93-151052
                                 931112
PAT. SEQ. LOC:
                  Disclosure; Fig 3A-3B
DATA ENTRY DATE:
                  25 APR 1997 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
                  97-020369 [02]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 97N-T58403
                  Human heat shock protein 65
DESCRIPTION:
KEYWORD:
                  Heat shock protein 65; Hsp65; insulin-dependent
                  diabetes mellitus; IDDM; autoimmune disease; diagnosis;
                  therapy; T cell; vaccine
ORGANISM:
                  Homo sapiens
ABSTRACT:
      The human heat shock protein 65 (Hsp65) (W01657) is expressed in
      the islets of the pancreas. The T cell response to Hsp65 is
      associated with the development of insulin-dependent diabetes
      mellitus (IDDM). A method for detecting the existence of, a
      tendency to develop, or the initiation of a process leading to IDDM
      involves detecting the presence of Hsp65 or antibodies or T cells
      reactive with the protein. Hsp65, when administered to a
      tolerogenic carrier, can be used to prevent or treat IDDM prior to
      development of clinical symptoms. Attenuated T cells can be used
      to vaccinate against autoimmunity to Hsp65 and to abort IDDM
AMINO ACID COUNTS:56 A; 22 R; 21 N; 35 D; 0 B; 3 C; 18 Q; 40 E; 0 Z;
                  58 G; 2 H; 41 I; 49 L; 54 K; 18 M; 11 F; 19 P; 24 S;
                  34 T; 1 W; 7 Y; 60 V;
SEQUENCE LENGTH:
                  573
SEQUENCE
        1 mlrlptvfrq mrpvsrvlap hltrayakdv kfgadaralm lqgvdllada
       51 vavtmgpkgr tviieqswgs pkvtkdgvtv aksid1kdky knigaklvqd
      101 vanntnegag dgtttatvla rsiakegfek iskganpvei rrgvmlavda
      151 viaetkkąsk pyttpeeiaą vatisangdk eigniisdam kkygrkgyit
      201 vkdgktlnde leiiegmkfd rgyispyfin tskgqkcefq dayvllsekk
```

101 vanntnegag dgtttatvla rsiakegfek iskganpvei rrgvmlavda
151 viaetkkqsk pvttpeeiaq vatisangdk eigniisdam kkvgrkgvit
201 vkdgktlnde leiiegmkfd rgyispyfin tskgqkcefq dayvllsekk
251 issiqsivpa leianlvlnr lkvglqvvav kapgflvlnr lkvglqvvav
301 kapgfgdnrk nq1kdmaiat ggavfgeeg1 t1n1edvqph dlgkvgeviv
351 tkddam11kg kgdkaqiekr iqeiieqldv ttseyekekl nerlak1sdg
401 vavlkvggts dvevnekkdr vtdalnatra aveegivlgg gcallrcipa
451 lds1tpaned qkigieiikr t1kipamtia knagvegs1i vekimqssse
501 vgydamagdf vnmvekgiid ptkvvrta1l daagvas1lt taevvvteip
551 keekdpgmga mggmgggmgg gmf

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

משת	ו זיירי	PF	ጥል	DI	æ.	٠

Key	Location Qualifier		
Peptide		Leader_peptide putative mitochondrial targeting sequence	
Protein	27573 label note	Mat_protein amino acid residues 266-285 differ from the translated	
Region		sequence (LVLNRLKVGLQVVAVKAPGF) "keratin-like region contg. Gly-Gly-Met repeats""	

```
L2
      ANSWER 5 OF 9 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 95P-R67385 Protein
                                            DGENE
                 DNA from Helicobacter pylori and Helicobacter felis -
TITLE:
                 used to develop prods. for detection, treatment and
                 prevention of Helicobacter infection
                 Ferrero R; Labigne A; Suerbaum S; Thiberge J
INVENTOR:
                  (INRM) INST NAT SANTE & RECH MEDICALE
PATENT ASSIGNEE:
                  INST PASTEUR
      (INSP)
PATENT INFO:
                 WO 9426901 A
                                 941124

    168 pp

APPLICATION INFO: WO 94-EP1625
                                 940519
PRIORITY INFO:
                 EP 93-401309
                                 930519
                                 931119
                 WO 93-EP3259
                 Disclosure; Fig. 7A(i-vii)
PAT. SEQ. LOC:
                 22 JUN 1995 (first entry)
DATA ENTRY DATE:
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
OTHER SOURCE:
                  95-006797 [01]
DESCRIPTION:
                 Mitochondrial protein Pl
KEYWORD:
                  Urease; immunogen; vaccine; diagnostic; heat shock
                  protein; HSP; GroEL-like protein; Helicobacter felis
ORGANISM:
                 Homo sapiens
ABSTRACT:
      The sequence of the Helicobacter pylori heat shock protein A (given
      in R67374) was compared to that of other GroEL-like proteins from
      Legionella pneumophila (R67381), Escherichia coli (R67382),
      Chlamydia psittaci (R67383), Mycobacterium leprae (R67384) and
      human mitochondrial protein Pl (R67385), and regions of homology
      were identified
AMINO ACID COUNTS:52 A; 17 R; 21 N; 37 D; 0 B; 3 C; 17 Q; 40 E; 0
                  56 G; 1 H; 41 I; 46 L; 52 K; 16 M; 10 F; 16 P; 26 S;
                  31 T; 1 W; 7 Y; 57 V;
SEQUENCE LENGTH:
                  547
SEQUENCE
        l ymadvkfgad aralmlqgvd lladavavtm gpkgrtviie qswgspkvtk
       51 dgvtvaksid 1kdkykniga klvqdvannt neeagdgttt atvlarsiak
      101 egfekiskga npveirrgvd lavdaviael kkqskpvttp eeiaqvatis
      151 angdkeigni isdamkkvgr kgvitvkdgk tlndeleiie gmkfdrgyis
      201 pyfintskgg kcefgdayvl lsekkissig sivpaleian lvlnrlkvgl
      251 qvvavkapgf lvlnrlkvgl qvvavkapgf gdnrknqlkd maistggsvf
      301 geegltlnle dvqphdlgkv gevivtkdda mllkgkgdka qiekriqeii
      351 eqldvttsey ekeklnerla klsdgvavlk vggtsdvevn ekkdrvtdal
      401 natraaveeg ivlgggcall rcipaldslt panedqkigi eiikrtlkip
      451 amtiaknagv dgslivekim qsssevgyda magdfvnmve kgiidptkvv
      501 rtalldaasv asllttaevv vteipeekdp gmgamggmgg gmgggmf
ALIGN Smith-Waterman score: 81
      38 aa overlap starting at 417
      csllicisllqlmvpvntdetieiivenkvkellanpa
      :::: :: :: ::: :: :: :: :: ::
```

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

callrcipaldsltpanedq_kigiei_ikrtlkipa

REFERENCE: Derwent DGene Search Report

FRANZ-BACON, et al., USSN: 09/099,898

Atty. Docket No.: DX0744K

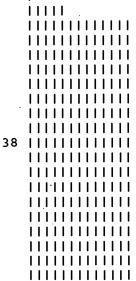
Mouse C2b

 ${\tt MKTTTCSLLICISLLQLMVPVNTEGTLESIVEKKVKELLANRDDCPSTVTKTFSCTSITASGRLASCPSGMTVTGCACGYGCGSWDIRDGNTCHCQCSTMDWATARCCQLA}$

13 ANSWERS FOUND ABOVE A THRESHOLD OF 67

Similarity Score

76 |



10

Answer Count

20

30

40

```
ANSWER 1 OF 13 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 96P-W05534 Peptide
                                           DGENE
                 New gClq receptor-based, HIV-1 gp 120 binding
TITLE:
                 peptide(s) - for preventing and treating HIV-1
                  infection
                 Fung M S C; Kim Y W; Sun B N V; Sun C R Y; Yu L
INVENTOR:
                  (TANO-N) TANOX BIOSYSTEMS INC
PATENT ASSIGNEE:
                 WO 9630400 Al 961003
                                                    53 pp
PATENT INFO:
                                960322
APPLICATION INFO: WO 96-US3905
                                950324
                 US 95-410360
PRIORITY INFO:
                 Disclosure; Page 46-48
PAT. SEQ. LOC:
DATA ENTRY DATE:
                  17 JAN 1997 (first entry)
DOCUMENT TYPE:
                  Patent
                  English
LANGUAGE:
                  96-455274 [45]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 96N-T41465
                  gClq receptor
DESCRIPTION:
                  gClq receptor; gClq-R; human immunodeficiency virus
KEYWORD:
                  type 1; HIV-1; gp120; immunogen; vaccine; therapy;
                  diagnosis
                  Homo sapiens
ORGANISM:
ABSTRACT:
      The gClq receptor (gClq-R) (W05534), a receptor for Clq complement,
      binds to HIV-1 gp120 and neutralises the infectivity of HIV-1.
      binding site for gpl20 has been identified (see also W05532).
      receptor exists on a variety of cell types, including B cells, T
      cells, monocytes, macrophages, neutrophils, eosinophils, platelets,
      fibroblasts and endothelial, liver, neural and smooth muscle cells.
      Recombinant, mature gClq-R can be produced in transformed host
      cells (see also T41465). It is useful for detecting or quantifying
      HIV-1 gpl20, HIV-1 virions or HIV-1 infected cells, and can also be
      used to treat or prevent HIV-1 infection and to raise antibodies of
      diagnostic or therapeutic value
AMINO ACID COUNTS:17 A; 15 R; 9 N; 20 D; 0 B; 7 C; 10 Q; 29 E; 0 Z;
                  20 G; 5 H; 9 I; 31 L; 16 K; 3 M; 13 F; 17 P; 20 S;
                  16 T; 3 W; 4 Y; 18 V;
SEQUENCE LENGTH: 282
SEQUENCE
        1 mlpllrcvpr vlgssvaglr aaapaspfrq llqpaprlct rpfgllsvra
       51 gserrpgllr prgpcacgcg cgslhtdgdk afvdflsdei keerkiqkhk
      101 tlpkmsggwe lelngteakl vrkvagekit vtfninnsip ptfdgeeeps
      151 qgqkveeqep eltstpnfvv eviknddgkk alvldchype devgqedeae
      201 sdifsirevs fqstgesewk dtnytlntds ldwalydhlm dfladrgvdn
      251 tfadelvels talehqeyit fledlksfvk sq
FEATURE TABLE:
               |Location|Qualifier|
|Pre-propeptide
               11..73
                        |label
Peptide
               |74..282 | label
                                  |Mat_protein
Protein
                                  |Glycosylation
                        |label
Modified_site | 114
                                  | "potential N-glycosylation
                        Inote
```

|site"

|site"

|Glycosylation

| *potential N-glycosylation

SEARCHED ON 26 OCT 1998

1136

Modified_site

|label

note

```
ANSWER 2 OF 13 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
                                           DGENE
ACCESSION NUMBER: 96P-R91446 Protein
                 Cloning of cDNA encoding cell surface antigen - useful
TITLE:
                 for isolation of diagnostic and therapeutic proteins
                 Aruffo A; Seed B
INVENTOR:
PATENT ASSIGNEE:
                 (GEHO)GEN HOSPITAL CORP
                                                    79 pp
PATENT INFO:
                 US 5506126 A 960409
APPLICATION INFO: US 88-160416
                                880225
PRIORITY INFO:
                 US 92-983647
                                921201
                 US 88-160416
                                880225
                                890713
                 US 89-379076
                                900713
                 US 90-553759
                 US 93-139273
                                931018
                 Example 16; Column 85-88
PAT. SEQ. LOC:
                 31 OCT 1996 (first entry)
DATA ENTRY DATE:
DOCUMENT TYPE:
                 Patent
LANGUAGE:
                 English
                 96-200279 [20]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 96N-T14726
                 Human CD53 antigen
DESCRIPTION:
                 Cell surface antigen; cloning; immunoselection;
KEYWORD:
                  immunotherapy; therapy; diagnosis; vector; COS; CD53;
                  lymphocyte
ORGANISM:
                 Homo sapiens
ABSTRACT:
     Human antigen CD53 (R91446) is a type III integral membrane protein
      that may be involved in the transport of factors essential for cell
     proliferation. Its amino acid sequence was deduced from a cDNA
      clone (T14726) obtd. using a novel immunoselection cloning
      technique. CD53 was expressed in transfected COS cells. Anti-CD53
      antibodies are a useful tool for the identification of
     haematopoietic neoplasms, and may prove helpful for identifying
     morphologically poorly defined cells
AMINO ACID COUNTS:10 A; 3 R; 11 N; 6 D; 0 B; 12 C; 5 Q; 6 E; 0 Z;
                  17 G; 5 H; 22 I; 33 L; 10 K; 6 M; 16 F; 4 P; 18 S;
                  10 T; 4 W; 7 Y; 14 V;
                 219
SEQUENCE LENGTH:
SEQUENCE
        1 mgmsslkllk yvlfffnllf wicgccilgf giyllihnnf gvlfhnlpsl
       51 tlgnvfvivg siimvvaflg cmgsikenkc llmsffilll iillaevtla
      101 illfvyegkl neyvakgltd sihryhsdns tkaawdsigs flqccgingt
      151 sdwtsgppas cpsdrkvegc yakarlwfhs nflyigiiti cvcvievlgm
      201 sfaltlncqi dktsqtigl
FEATURE TABLE:
               |Location|Qualifier|
Key
|Hydrophobic_domain
               18..36
Domain
                        |label
                                  |Hydrophobic_domain
               155..75
                       |label
Domain
                                  [Hydrophobic_domain
               |81..106 | label
Domain
Modified_site |149..151|label
                                  |Glycosylation_site
Modified_site |228..230|label
                                  [Glycosylation_site
```

|Hydrophobic_domain

SEARCHED ON 26 OCT 1998

Domain

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

|182..206|1abel

ALIGN Smith-Waterman score: 73
96 aa overlap starting at 75
mktttcsll_icisllqlmvpvntegtlesivekkvkella_____nrddcp
.: .: .: .: .: .: .: .: .:
ikenkc1lmsffillliillaevtlai11fvyeqklneyvakgltdsihryhsdnstkaa
_stvtktfsctsi__t_asgrlascpsgmtvtgc
... .: :: :: :: ::
wdsiqsflqccgingtsdwtsgppascpsdrkvegc

```
ANSWER 3 OF 13 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
                                            DGENE
ACCESSION NUMBER: 94P-R63790 Protein
                 New xylanase enzymes from Aspergillus aculeatus - used
TITLE:
                  for degrading plant cell wall components, e.g. in the
                  prepn. of feed, in baking and in prepn. of pulp or
                  paper
                  Andersen L N; Christgau S; Dalboge H; Heldt-hansen H P;
INVENTOR:
                  Jacobsents; Kauppinen M S; Kofod L V; Mullertz A; Munk
                  N; Si J Q
                  (NOVO) NOVO-NORDISK AS
PATENT ASSIGNEE:
PATENT INFO:
                  WO 9421785 A 940929
                                                     80 pp
APPLICATION INFO: WO 94-DK88
                                 940302
PRIORITY INFO:
                  DK 93-268
                                 930310
                  DK 93-1151
                                 931014
                  Claim 10; Page 57
PAT. SEQ. LOC:
DATA ENTRY DATE:
                  07 JUN 1995 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
OTHER SOURCE:
                  94-317006 [39]
CROSS REFERENCES: N-PSDB: 94N-Q74637
DESCRIPTION:
                  Aspergillus aculeatus xylanase II
                  Xylanase II; Aspergillus aculeatus;
KEYWORD:
                  alpha-arbino-pyranosidase; brewing; paper pulp; food
                  preparation; plant cell wall degradation
ORGANISM:
                  Aspergillus aculeatus
ABSTRACT:
      Q74637 encodes R63790 Aspergillus aculeatus xylanase II, which
      degrades plant cell wall components and reduces the viscosity of
      plant cell wall derived material. These properties are useful in
      the production of dough and baked products; in the preparation of
      feed, food, beer, wine, pulp and paper; and for the separation of
      cereal components. In addition xylanase II exhibits
      alpha-arbino-pyranosidase activity, and it can also be used in the
      production of antibodies
AMINO ACID COUNTS:54 A; 6 R; 22 N; 21 D; 0 B; 8 C; 20 Q; 10 E; 0 Z;
                  39 G; 9 H; 17 I; 38 L; 18 K; 4 M; 13 F; 16 P; 28 S;
                  64 T; 11 W; 20 Y; 33 V; 2 Others;
SEQUENCE LENGTH:
                  453
SEQUENCE
        1 mvgllsitaa laatvlpniv savgldqaav akglqyfgta tdnpeltdip
       51 yvtq1nntad fgqitpgnsm kwdatepsqg tftftkgdvi adlaegngqy
      101 lrchtlvwyn qlpswvtsgt wtnatltaal knhitnvvsh ykgkclhwdv
      151 vnealnddgt yrtnifytti geayipiafa aaaaadpdak lfyndynley
      201 ggakaasara ivqlvknaga kidgvglqah fsvgtvpsts slvsvlqsft
      251 algvevayte advrillptt attlaqqssd fqalvqscvq ttgcvgftiw
      301 dwtdkyswvp stfsgygaal pwdenlvkkp ayngllagmg vtvtttttt
      351 tatatgkttt tttgatstgt taahwgqcgg lnwsgptaca tgytctyvnd
      401 yysqclxsia qpkpagvlai qsvrfiyhnt qnslldlxkk ktlehtggrs
      451 smh
```

```
FEATURE TABLE:
```

SEARCHED ON 26 OCT 1998

SEARCHED ON 26 OCT 1998 FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

ANSWER 4 OF 13 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 93P-R45352 protein DGENE

TITLE: New polypeptide(s) from venom of Plectreurys tristis -

used as insecticidal agents which are paralytic and/or

toxic to insects at low concns

INVENTOR: Leisy D J; Quistad G B; Skinner W S

PATENT ASSIGNEE: (SANO) SANDOZ AG

(SANO) SANDOZ PATENT GMBH

(SANO) SANDOZ-ERFINDUNGEN VERW GES MBH

PATENT INFO: EP 556160 A 930818 50 pp

APPLICATION INFO: EP 93-810078 930208 PRIORITY INFO: US 92-837194 920211

PAT. SEQ. LOC: Claim 1; Page 42

DATA ENTRY DATE: 04 FEB 1994 (first entry)

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 93-260398 [33]

DESCRIPTION: Arachnoidal plectoxin polypeptide G

KEYWORD: Primitive Hunting Spider; insecticidal; plectoxin;

arachnid

ORGANISM: Plectreurys tristis

ABSTRACT:

This polypeptide corresponds to a plectoxin sequence isolated from the Primitive Hunting Spider (Plectreurys tristis). The plectoxins have insecticidal properties and are used against Lepidopteran species. See also R38840 and R45351

AMINO ACID COUNTS:3 A; 2 R; 2 N; 1 D; 0 B; 10 C; 1 Q; 1 E; 0 Z; 7

G; 0 H; 1 I; 3 L; 7 K; 0 M; 2 F; 0 P; 5 S; 1

T; 1 W; 0 Y; 2 V;

SEQUENCE LENGTH: 49

SEQUENCE

l gckgflvkcd snseccktai vkgkkkqlsc lcgawgagcs csfrcgnrc

ALIGN Smith-Waterman score: 76

32 aa overlap starting at 16 c_tsitasg_rlascpsgmtvtgcacgygcgs

cktaivkgkkkglsclcgawgagcscsfrcgn

```
ANSWER 5 OF 13 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 92P-R20818 Protein
                                            DGENE
                  New CD53 cell surface antigen and DNA encoding it - for
TITLE:
                  immuno-therapy and diagnosis of haematopoietic
                  neoplasms, etc
                  Seed B; Aruffo A; Amiot M
INVENTOR:
                  (GEHO-N)GEN HOSPITAL CORP
PATENT ASSIGNEE:
PATENT INFO:
                  WO 9201049 A 920123
                                                    160 pp
APPLICATION INFO: WO 90-US4986
                                 900715
PRIORITY INFO:
                  US 90-553759
                                 900713
                  Claim 4; Page 123
PAT. SEQ. LOC:
                  21 MAY 1992 (first entry)
DATA ENTRY DATE:
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
OTHER SOURCE:
                  92-056864 [07]
CROSS REFERENCES: N-PSDB: 92N-Q21187
                  CD53 haematopoietic antigen
DESCRIPTION:
KEYWORD:
                  Rapid immunoselection cloning technique; cell surface
                  antigen; haematopoietic neoplasm; type III integral
                  membrane protein
ORGANISM:
                  Homo sapiens
ABSTRACT:
      A cDNA clone encoding CD53 was obtained using the rapid
      immunoselection cloning method (see e.g Q21164 for description of
      method). The cDNA libraries were prepared from the promyelocytic
      tumour cell line HL60 and from peripheral blood lymphocytes and
      transfected into COS cells. The first of the four predicted
      hydrophobic regions is atypically long for either a signal sequence
      or a simple transmembrane alpha-helix. Both cysteine and glycine
      have been found to precede the signal cleavage site (Von Heijne,
      Nucleic Acid Res. 14:4683 (1986)) and the presence of 3 cysteines
      and a glycine in the middle of the first hydrophobic region
      suggests that the N-terminus of the mature CD53 protein begins
      there
AMINO ACID COUNTS: 10 A; 3 R; 11 N; 6 D; 0 B; 12 C; 5 Q; 6 E; 0 Z;
                  17 G; 5 H; 22 I; 33 L; 10 K; 6 M; 16 F; 4 P; 18 S;
                  10 T; 4 W; 7 Y; 14 V;
SEQUENCE LENGTH:
                  219
SEQUENCE
        1 mgmsslkllk yvlfffnllf wicgccilgf giyllihnnf gvlfhnlpsl
       51 tlgnvfvivg siimvvaflg cmgsikenkc llmsffilll iillaevtla
      101 illfvyeqkl neyvakgltd sihryhsdns tkaawdsiqs flqccgingt
      151 sdwtsgppas cpsdrkvegc yakarlwfhs nflyigiiti cvcvievlgm
```

FEATURE TABLE:

Key	Location Qualifier		
Region	=+====================================	[hydrophobic	
Region	5575 label	Ihydrophobic	
Region	81106 label	hydrophobic	
Region	182206 label	[hydrophobic	
Modified_site	149151 1abel note	N-linked_glycosylation "putative"	
Modified_site	168170 1abel	N-linked_glycosylation	
	Inote	"putative"	

SEARCHED ON 26 OCT 1998

201 sfaltlncqi dktsqtigl

ALIGN Smith-Waterman score: 73
96 aa overlap starting at 75
mktttcsll_icisllqlmvpvntegtlesivekkvkella_____nrddcp
.: .: :: :: :: :: :: :: :: :: ::
ikenkcllmsffillliillaevtlaillfvyeqklneyvakgltdsihryhsdnstkaa
_stvtktfsctsi___t_asgrlascpsgmtvtgc
... :: :: :: :: :: :: :: ::
wdsiqsflqccgingtsdwtsgppascpsdrkvegc

REFERENCE: Derwent DGene Search Report

50

40

FRANZ-BACON, et al., USSN: 09/099,898

Atty. Docket No.: DX0744K

Human C10

 ${\tt MGPSSCLLLILIPLLQLINPGSTQCSLDSVMDKKIKDVLNSLEYSPSPISKKLSCASVKSQGRPSSCPAGMAVTGCACGYGCGSWDVQLETTCHCQCSVVDWTTARCCHLT}$

30

25 ANSWERS FOUND ABOVE A THRESHOLD OF 67

Similarity Score ' 94 | | | 11111 IIIIII \mathbf{I} 1111111111111111111111 1111111111111111111111111111111 11111111111111111111111111111 11111111111111111111111111111 111111111111111111111111111111 1111111111111111111111111111 11111111111111111111111111111111 11111111111111111111111111111 1111111111111111111111111111111 111111111111111111111111111111 Answer Count 10 20

SEARCHED ON 26 OCT 1998 FILE LAST UPDATED: 18 OCT 1998 <19981018/UP> ANSWER 1 OF 25 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 98P-W60854 Protein DGENE

New bovine polypeptide that activates mammalian B TITLE:

cell(s) - used e.g. to treat T cell immunodeficiency or allergy, as vaccine adjuvant, as T cell surrogate for infants, and for monoclonal antibody production, also specific antibodies for treating B cell hyperactivity

Alizadeh-Khiavi K; Filipp D; Julius M H

INVENTOR: (WELL-N)WELLESLEY HOSPITAL FOUND

PATENT ASSIGNEE: WO 9822580 A2 980528 64 pp PATENT INFO:

APPLICATION INFO: WO 97-CA880 971118 US 96-746883 961118 PRIORITY INFO:

Claim 14; Fig 7 PAT. SEQ. LOC:

01 OCT 1998 (first entry) DATA ENTRY DATE:

DOCUMENT TYPE: Patent LANGUAGE: English

98-312466 [27] OTHER SOURCE: CROSS REFERENCES: N-PSDB: 98N-V37228

Human CD14 protein DESCRIPTION: CD14; B cell activator; bovine lactation-associated KEYWORD:

immunotropic protein; LAIT; T cell immunodeficiency; X-linked hyper-IgM syndrome; allergy; common variable

immunodeficiency; X-linked agammaglobulinaemia;

vaccine; infant feeding formulae; human

Homo sapiens ORGANISM:

ABSTRACT:

This sequence is the human CD14 protein of the invention. The CD14 protein was used to isolate the bovine CD14 of the invention, which is able to activate mammalian B cells. The protein is also known as bovine lactation-associated immunotropic protein (LAIT), and is used to activate B cells, particularly in humans. Particularly it is administered to subjects: (a) with CD40 negative or deficient B cells; (b) suffering from T cell immunodeficiency (e.g. X-linked hyper-IgM syndrome, common variable immunodeficiency or X-linked agammaglobulinaemia) or allergy (i.e. with CD40 ligand negative or defective T cells); or (c) to induce growth and differentiation of B cells to highly productive Ig secreting cells. Particular applications are in infant feeding formulae (as immunostimulant) and as adjuvant in vaccines (optionally with bovine CD14 coupled to the antigen). The DNA sequences are also used to enrich mammalian B cells secreting a monoclonal antibody (MAb) of particular antigenic specificity, by activating cells with sub-optimal amount of the DNA in combination with the antigen. The enriched B cells are then used to produce hybridomas that produce specific MAb. Antibodies raised against human CD14 are used to reduce/inhibit activity of B cells that are hyperactivated by high serum levels of CD14. Bovine CD14 stimulates growth (induce DNA synthesis) in resting murine spleen cells and is 200 times more effective than lipopolysaccharide (LPS), with the effect unaffected by presence of serum. It also induces Ig secretion and a partial isotype switch from IgM to IgG, in absence of T cells

AMINO ACID COUNTS:43 A; 22 R; 15 N; 16 D; 0 B; 10 C; 14 Q; 20 E; 0 Z;

24 G; 7 H; 4 I; 62 L; 9 K; 6 M; 10 F; 30 P; 29 S;

18 T; 5 W; 3 Y; 28 V;

SEQUENCE LENGTH: 375

SEQUENCE 1 merascllll llplvhvsat tpepceldde dfrcvcnfse pqpdwseafq 51 cvsaveveih agglnlepfl krvdadadpr qyadtvkalr vrrltvgaaq 101 vpaqllvgal rvlaysrlke ltledlkitg tmpplpleat glalsslrlr 151 nvswatgrsw laelqqwlkp glkvlsiaqa hspafsyeqv rafpaltsld 201 lsdnpglger glmaalcphk fpaiqnlalr ntgmetptgv caalaaagvq 251 phsldlshns lratvnpsap rcmwssalns lnlsfagleq vpkglpaklr 301 vldlscnrln rapqpdelpe vdnltldgnp flvpgtalph egsmnsgvvp 351 acarstlsvg vsgtlvllqg argfa FEATURE TABLE: |Location|Qualifier| Key | encoded by TGC Misc_difference |87 note ALIGN Smith-Waterman score: 94 58 aa overlap starting at 1 ${\tt mgpssclllilipllqlinpgstqcsldsvmdkkikdvlnsleyspspiskklscasv}$

```
ANSWER 2 OF 25 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
                                            DGENE
ACCESSION NUMBER: 98P-W41693 Protein
                  Assay for lipo:poly:saccharide binding inhibitors -
TITLE:
                 useful in the treatment of septic shock and other
                  lipo:poly:saccharide-mediated disorder(s)
                 Mintz D N; Tobias P; Ulevitch R
INVENTOR:
                  (SCRI)SCRIPPS RES INST
PATENT ASSIGNEE:
PATENT INFO:
                 US 5705398 A 980106
                                                     21 pp
APPLICATION INFO: US 94-205719
                                 940302
PRIORITY INFO: \
                 US 94-205719
                                 940302
                 Claim 9; Columns 19-22
PAT. SEQ. LOC:
                  01 MAY 1998 (first entry)
DATA ENTRY DATE:
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
OTHER SOURCE:
                  98-086145 [08]
CROSS REFERENCES: N-PSDB: 98N-V05505
                  Human CD14
DESCRIPTION:
                  Identification; binding inhibitor; lipopolysaccharide;
KEYWORD:
                  LPS; CD14; binding protein; LBP; monocyte receptor;
                  treatment; septic shock; human
ORGANISM:
                  Homo sapiens
ABSTRACT:
      The present sequence was used in the development of a novel method
      for identifying a compound that inhibits the binding of a
      lipopolysaccharide (LPS) to a LPS-binding protein (LBP), or
      LBP-dependent binding of LPS to monocyte receptor CD14. The method
      comprises measuring the fluorescence emitted by a reaction mixture
      containing fluoresceinated LPS, isolated LBP and optionally CD14 in
      the presence and absence of the compound, and identifying the
      compound as an inhibitor if the fluorescence emitted by the
      reaction mixture containing the compound is less than that emitted
      by the reaction mixture that does not contain the compound. The
      method can be used to identify drugs useful for treating septic
      shock and related LPS-mediated disorders
AMINO ACID COUNTS:42 A; 23 R; 15 N; 16 D; 0 B; 11 C; 14 Q; 20 E; 0 Z;
                  24 G; 7 H; 4 I; 62 L; 9 K; 6 M; 10 F; 30 P; 29 S;
                  18 T; 5 W; 2 Y; 28 V;
SEQUENCE LENGTH:
                  375
SEQUENCE
        l merascilli lipivhvsat tpepceldde dfrcvcnfse pqpdwseafq
       51 cvsaveveih agglnlepfl krvdadrdpr qyadtvkalr vrrltvgaaq
      101 vpaq11vgal rvlaysr1ke ltledlkitg tmpplpleat glalsslrlr
      151 nvswatgrsw lae1qqwlkp glkvlsiaqa hspafsceqv rafpaltsld
      201 lsdnpglger glmaalcphk fpaiqnla1r ntgmetptgv caalaaagvq
      251 phs1d1shns lratvnpsap rcmwssalns lnlsfagleq vpkglpaklr
      301 vldlscnrln rapqpdelpe vdnltldgnp flvpgtalph egsmnsgvvp
      351 acarstlsvg vsgtlvllgg argfa
ALIGN Smith-Waterman score: 94
      58 aa overlap starting at 1
      mgpssc111i1ip11q1inpgstqcs1dsvmdkkikdv1ns1eyspspiskk1scasv
                          : :: :. .. : : ... : .......
     : .:::::::::::
```

merascllllllplvhvsattpepceld___dedfrcvcnfsepqpd_wseafqcvsa

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

```
ANSWER 3 OF 25 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 96P-W05316 Protein
                 Recombinant DNA encoding myelomonocytic differentiation
TITLE:
                  antigen CD14 - used for producing recombinant CD14 for
                 use in e.g. diagnosis of myeloid disorders such as
                  leukaemia
                 Goyert S M
INVENTOR:
                  (GOYE-I)GOYERT S M
PATENT ASSIGNEE:
                 US 5543303 A 960806
PATENT INFO:
                                                    11 pp
                                881128
APPLICATION INFO: US 88-276794
PRIORITY INFO:
                 US 88-276794
                                 881128
                 US 90-536163
                                900608
                 US 91-787763
                                911106
                 US 92-916806
                                920722
                  US 93-165583
                                931213
PAT. SEQ. LOC:
                 Claim 1; Fig 3
DATA ENTRY DATE:
                  03 JAN 1997 (first entry)
DOCUMENT TYPE:
                  Patent
                  English
LANGUAGE:
                  96-370638 [37]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 96N-T39716; N-PSDB; 96N-T39717
                 Myelomonocytic differentiation antigen CD14
DESCRIPTION:
                 Myelomonocytic differentiation antigen; CD14; myeloid
KEYWORD:
                  leukaemia; diagnosis
ORGANISM:
                  Homo sapiens
ABSTRACT:
      Human myelomonocytic differentiation antigen CD14 (W05316) is an
      antigen useful in the diagnosis of mature myeloid leukemia. Its
      amino acid sequence was deduced from a cDNA clone (T39717) obtd. by
      screening COS 7 cell transfectants with monoclonal antibodies to
      CD14. Large amts. of CD14 can be produced by expression in
      transformed host cells; mature, glycosylated CD14 is produced in
      mammalian host cells, and nonglycosylated CD14 in prokaryotic hosts
AMINO ACID COUNTS:43 A; 22 R; 15 N; 16 D; 0 B; 11 C; 14 Q; 20 E; 0 Z;
                  24 G; 7 H; 4 I; 62 L; 9 K; 6 M; 10 F; 30 P; 29 S;
                  18 T; 5 W; 2 Y; 28 V;
                  375
SEQUENCE LENGTH:
SEQUENCE
        1 merascllll llplvhvsat tpepceldde dfrcvcnfse pqpdwseafq
       51 cvsaveveih agglnlepfl krvdadadpr qyadtvkalr vrrltvgaaq
      101 vpaqllvgal rvlaysrlke ltledlkitg tmpplpleat glalsslrlr
      151 nvswatgrsw laelqqwlkp glkvlsiaqa hspafsceqv rafpaltsld
      201 lsdnpglger glmaalcphk fpaignlalr ntgmetptgv caalaaagvq
      251 phsldlshns lratvnpsap rcmwssalns lnlsfagleq vpkglpaklr
      301 vldlscnrln rapqpdelpe vdnltldgnp flvpgtalph egsmnsgvvp
      351 acarstlsvg vsgtlvllqg argfa
FEATURE TABLE:
               |Location|Qualifier|
Key
______+
               11..19
                        |label
                                  |Sig_peptide
Modified_site
               137..39
                        |label
                                  |Glycosylation
                                  | potential N-linked
                        Inote
                                  |glycosylation site"
```

|Glycosylation

| potential N-linked

SEARCHED ON 26 OCT 1998

Modified_site

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

|151..153|label

Inote

		•
~	1 1	glycosylation site"
Modified_site	266268 label	Glycosylation
•	Inote	"potential N-linked
	1	glycosylation site"
Modified_site	282284 label	Glycosylation
•	Inote	*potential N-linked
	1	glycosylation site"
Modified_site	323325 label	Glycosylation
	note	*potential N-linked
	1 1 .	glycosylation site"

ANSWER 4 OF 25 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 95P-R71730 Protein DGENE.

New merosin fragments, corresp. DNA and antibodies -TITLE:

for diagnosing tumour malignancy, promoting or

inhibiting neurite growth and promoting cell attachment

INVENTOR: Engvall E; Leivo I

(LJOL-N)LA JOLLA CANCER RES FOUND PATENT ASSIGNEE:

WO 9508628 A2 950330 65 pp PATENT INFO:

APPLICATION INFO: WO 94-US10730 940921 PRIORITY INFO: US 93-125077 930922

Claim 5; Fig 6 PAT. SEQ. LOC:

01 MAY 1996 (first entry) DATA ENTRY DATE:

DOCUMENT TYPE: Patent LANGUAGE: English

95-139597 [18] OTHER SOURCE:

CROSS REFERENCES: N-PSDB: 95N-Q86480 AND T17419

DESCRIPTION: Merosin major subunit

Human; 380-400 kD; merosin; major subunit; placenta; KEYWORD:

straited muscle; peripheral nerve; trophoblast; Schwann cell neoplasm; 65 kD subunit; 80 kD subunit; merosin polypeptide; merosin subunit; M chain; laminin M chain;

antigen; antibody; detection; tumour; malignancy;

neurite outgrowth; inhibitor; cell attachment

ORGANISM: Homo sapiens

ABSTRACT:

This sequence represents the human 380-400 kD merosin major subunit. Merosin is an isoform of laminin and shows structural and sequence similarity to the human laminin A chain. Mature human merosin is 30 amino acids larger than the human laminin A chain. Similarly to all laminin chains, the merosin protein has distinct domains which are predicted to have globular regions, cysteine-rich rod-like regions and helical structures (see features table). Merosin has a large globular domain at the carboxy terminal end. The merosin protein has an apparent mol. wt. of about 800 kD and is composed of four polypeptides with molecular weights of 300, 200, 200 and 80 kD. The 300 kD polypeptide is joined to the 200 kD polypeptides by disulphide bonds and the 300 and 80 kD polypeptides comprise the major subunit protein given in R71729. Merosin is found in placenta, straited muscle, peripheral nerve, trophoblasts and human Schwann cell neoplasms, amoung other tissues. 380-400 major merosin subunit also yields a 65 kD subunit. The 380-400 merosin subunit has been designated merosin polypeptide, merosin subunit, M chain or laminin M chain. Fragments of the merosin protein may be used as antigens to raise anti-merosin antibodies. These antibodies may be used in the detection of merosin, as the absence of merosin in a tumour sample indicates malignancy. Contacting a neurone with merosin promotes neurite outgrowth. The merosin polypeptide may also be used in contacting inhibitors of neurite outgrowth, thereby also promoting the outgrowth. Merosin also promotes cell attachment. The merosin gene has been mapped to chromosome 6, more precisely to bands 6a22->a23

AMINO ACID COUNTS: 205A; 159R; 162N; 183D; 0 B; 162C; 119Q; 202E; 0

261G; 71 H; 166I; 246L; 184K; 46 M; 103F; 173P; 194S;

193T; 29 W; 96 Y; 156V;

SEQUENCE LENGTH: 3110

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

SEQUENCE

1 mpgaagvl11 ll1sgglggv qaqrpqqqrq sqahqqrglf pavlnlasna 51 littnatcge kgpemycklv ehvpgqpvrn pqcricnqns snpnqrhpit 101 naidgkntww qspsikngie yhyvtitldl qqvfqiayvi vkaansprpg 151 nwilersldd veykpwqyha vtdtec1tly niyprtgpps yakddevict 201 sfyskihple ngeihislin grpsaddpsp elleftsary irlrfqrirt 251 lnadlmmfah kdpreidpiv trryyysvkd isvggmcicy gharacpldp 301 atnksrcece hntcgdscdq ccpgfhqkpw ragtfltkte ceacnchgka 351 eecyydenva rrnlslnirg kyigggvcin ctqntaginc etctdgffrp 401 kgvspnyprp cqpchcdpig slnevcvkde kharrglapg schcktgfgg 451 vscdrcargy tgypdckacn csglgskned pcfgpcicke nveggdcsrc 501 ksgffnlqed nwkgcdecfc sgvsnrcqss ywtygkiqdm sgwyltdlpg 551 rirvapqqdd 1dspqqisis naearqalph syywsapapy 1gnklpavgg 601 qltftisydl eeeeedterv lqlmiilegn dlsistaqde vylhpseeht 651 nvlllkeesf tihgthfpvr rkefmtvlan lkrvllqity sfgmdaifrl 701 ssvnlesavs yptdgsiaaa vevcqcppgy tgsscescwp rhrrvngtif 751 ggicepcqcf ghaescddvt geclnckdht ggpycdkclp gfygeptkgt 801 sedcqpcacp lnipsnnfsp tchldrslgl icdgcpvgyt gprcercaeg 851 yfgqpsvpgg scqpcqcndn ldfsipgscd slsgsclick pgttgrycel 901 cadgyfgdav dakncqpcrc naggsfsevc hsqtgqcecr anvqgqrcdk 951 ckagtfglqs argcvpcncn sfgsksfdce esgqcwcqpg vtgkkcdrca 1001 hgyfnfqegg ctacecshlg nncdpktgrc icppntigek cskcapntwg 1051 hsittgckac ncstvgsldf qcnvntgqcn chpkfsgakc tecsrghwny 1101 prcnlcdcfl pgtdattcds etkkcscsdq tgqctckvnv egihcdrcrp 1151 gkfgldaknp lgcsscycfg tttqcseakg lirtwvtlka eqtilplvde 1201 alghtttkgi vfghpeivah mdlmredlhl epfywklpeg fegkklmayg 1251 gklkyaiyfe areetgfsty npqviirggt pthariivrh maapligqlt 1301 rheiemteke wkyygddprv hrtvtredf1 dilydihyil ikatygnfmr 1351 qsriseisme vaeqgrgttm tppadliekc dcplgysgls ceaclpgfyr 1401 lrsqpggrtp gptlgtcvpc qcnghss1cd petsicqncq hhtagdfcer 1451 calgyygivk glpndcqqca cplisssnnf spscvaegld dyrctacprg 1501 yegqycerca pgytgspgnp ggscqececd pygslpvpcd pvtgfctcrp 1551 gatgrkcdgc khwharegwe cvfcgdectg lllgdlarle qmvmsinltg 1601 plpapykmly glenmtqelk hllspqrape rliqlaegnl ntlvtemnel 1651 ltratkvtad geqtgqdaer tntrakslge fikelardae avnekaikln 1701 etlgtrdeaf ernleglqke idqmikelrr knletqkeia edelvaaeal 1751 lkkvkklfge srgeneemek dlrekladyk nkvddawdll reatdkirea 1801 nrlfavngkn mtalekkkea vesgkrgien tlkegndild eanrladein 1851 siidyvedig tklppmseel ndkiddlsge ikdrklaekv sqaeshaaql 1901 ndssavldgi ldeaknisfn ataafkaysn ikdyideaek vakeakdlah 1951 eatklatgpr gllkedakgc lqksfrilne akklandvke nedhlnglkt 2001 rienadarng dllrtlndtl gklsaipndt aaklqavkdk arqandtakd 2051 vlaqitelhq nldg1kknyn kladsvaktn avvkdpsknk iiadadatvk 2101 nleqeadrli dklkpikele dnlkknisei kelinqarkq ansikvsvss 2151 ggdcirtykp eikkgsynni vvnvktavad nllfylgsak fidflaiemr 2201 kgkvsflwdv gsgvgrveyp dltiddsywy rivasrtgrn gtisvraldg 2251 pkasivpsth hstsppgyti 1dvdanamlf vggltgklkk adavrvitft 2301 gcmgetyfdn kpiglwnfre kegdckgctv spqvedsegt iqfdgegyal 2351 vsrpirwypn istvmfkfrt fsssallmyl atrdlrdfms veltdghikv 2401 sydlgsgmas vvsnqnhndg kwksftlsri qkqanisivd idtnqeenia 2451 tsssgnnfgl dlkaddkiyf gglptlrnls mkarpevnlk kysgclkdie 2501 isrtpynils spdyvgvtkg cslenvytvs fpkpgfvels pvpidvgtei 2551 nlsfstknes giillgsggt papprrkrrq tgqayyvill nrgrlevhls 2601 tgartmrkiv irpepnlfhd grehsvhver trgiftvqvd enrrymqnlt 2651 veqpievkkl fvggappefq psplrnippf egciwnlvin svpmdfarpv 2701 sfknadigrc ahqklreded gaapaeiviq pepvptpafp tptpvlthgp 2751 caaesepall igskqfglsr nshiaiafdd tkvknrltie levrteaesg 2801 llfymaainh adfatvqlrn glpyfsydlg sgdthtmipt kindgqwhki

2851 kimrskqegi lyvdgasnrt ispkkadild vvgmlyvggl pinyttrrig 2901 pvtysidgcv rnlhmaeapa dleqptssfh vgtcfanaqr gtyfdgtgfa 2951 kavggfkvgl dllvefefat ttttgvllgi ssqkmdgmgi emideklmfh 3001 vdngagrfta vydagvpghl cdgqwhkvta nkikhrielt vdgnqveaqs 3051 pnpastsadt ndpvfvggfp ddlkqfgltt sipfrgcirs lkltkgtash 3101 wrlilprpwn

FEATURE TABLE: '

Key	-	Qualifier	l
Region		•	region encoded by Q86480"
			"Signal peptide"
Domain	•		Domain VI
•		Inote	*predicted to form globular
	İ		structure"
Modified_site	15557	Inote	"N-linked glycosylation site"
Modified_site			"N-linked glycosylation site"
Domain	1287527	label	Domain V
	I	note	"contains four and one half
	1		Cystein-rich EGF-like repeats,
•	1		predicted to have rigid
	1	•	rod-like structure"
Modified_site	1303305		"N-linked glycosylation site"
Modified_site	• • • • •		"N-linked glycosylation site"
Modified_site	•		"N-linked glycosylation site"
Modified_site	•	•	"N-linked glycosylation site"
Domain	1528723	label	Domain IVb
	1	Inote	predicted to form globular
	1	I	structure"
Domain	17241175	•	Domain IIIb
	I	Inote	"contains ten and one half
	!	!	Cystein-rich EGF-like repeats,
	1	!	predicted to have rigid
M. 1151 1 . 11		l .	rod-like structure"
Modified_site		Inote	"N-linked glycosylation site"
Modified_site	10611063		"N-linked glycosylation site"
Domain	11761379		Domain IVa
		note	"predicted to form globular structure "
Domain	111801573	l llabol	Domain IIIa
DOMATH	11100,.1373	Inote	"contains four Cystein-rich
	1	I	EGF-like repeats, predicted
	1	! 	to have rigid rod-like
	1	1	Istructure"
Domain	15742153	ı Ilahol	Domain I+II
Domain	1	Inote	"forms two B-type chains,
	i .	1	forms triple coiled-coil
	i	i i	Istructure"
Modified_site	15971599	Inote	"N-linked glycosylation site"
Modified_site	116141616		"N-linked glycosylation site"
Modified_site	117001702		"N-linked glycosylation site"
Modified_site	118101812		"N-linked glycosylation site"
Modified_site	119011903		"N-linked glycosylation site"
Modified_site	19161918		"N-linked glycosylation site"
Modified_site	119201922		"N-linked glycosylation site"
Modified_site	120172019	note	"N-linked glycosylation site"

```
| "N-linked glycosylation site"
Modified_site
               |2028...2030|note
                                     | "N-linked glycosylation site"
Modified_site
               |2045..2047|note
                                     |"N-linked glycosylation site"
               |2126..2128|note
Modified_site
                                     | Domain G
Domain
               |2154..3110|label
                                     |"forms large globule at end of
                           Inote
                                     |laminin long arm"
                                     |"N-linked glycosylation site"
               |2240..2242|note
Modified_site
                                     |"N-linked glycosylation site"
Modified_site
               |2360..2362|note
                                     | "N-linked glycosylation site"
Modified_site
               |2435..2437|note
                                     |"N-linked glycosylation site"
Modified_site
               |2478..2480|note
                                     |"N-linked glycosylation site"
               |2551..2553|note
Modified_site
                                     |"N-linked glycosylation site"
Modified_site
               |2558..2560|note
                                     | "N-linked glycosylation site"
Modified_site
               |2648..2650|note
                                     |"N-linked glycosylation site"
Modified_site |2868..2870|note
                                        | "N-linked glycosylation site"
   Modified_site
                 |2893..2895|note
ALIGN Smith-Waterman score: 97
      64 aa overlap starting at 939
      c_asvksqgrpsscpag___mavtgca___c_gygcgswdvqlettchcqcsvvdwtta
                           ... ::.
      : ::::: : :::
                                    : ..: :.: . : :: .:.
      cranvqgq_rcdkckagtfglqsargcvpcncnsfgsksfdceesgqcwcqpgvtgkkcd
      rcch
      ::::
      rcah
```

```
ANSWER 5 OF 25 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 90P-R07450 protein
                                          DGENE
TITLE:
                 DNA encoding TNF binding protein and TNF- receptor -
                 used in tumour treatment and to understand mechanismsm
                 to TNF action
INVENTOR:
                 Hauptmann R; Himmler A; Maurer-Fogy I; Stratowa C
PATENT ASSIGNEE:
                 (BOEH) BOEHRINGER INGELHEIMINT
PATENT INFO:
                 EP 393438
                           A 901024
                                                   51 pp
APPLICATION INFO: EP 90-106624
                                900406
PRIORITY INFO:
                 DE 89-3920282 890621
                 DE 89-3913101 890421
                 Disclosure; Fig 8(1-2)
PAT. SEQ. LOC:
DATA ENTRY DATE:
                 29 JAN 1991 (first entry)
DOCUMENT TYPE:
                 Patent -
LANGUAGE:
                 German -
                 90-321987 [43]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 90N-Q06284
                 Rat Tumour Necrosis Factor-Receptor from raTNF-R8 cDNA
DESCRIPTION:
                 Tumour necrosis factor binding protein; TNF-BP;
KEYWORD:
                 TNF-receptor; raTNF-R8
ORGANISM:
                 Rat rattus
ABSTRACT:
     A rat brain cDNA analogue of the HS913T cDNA library from rat
     glioma cell line C6 (ATCC CCL107) is prepared in lambda-gt11. The
      isolated clone raTNF-R8 is used as probe to isolated the entire
     human TNF receptor, as represented in Q06285. See also
     Q06282-Q06285
AMINO ACID COUNTS: 22 A; 26 R; 20 N; 16 D; 0 B; 31 C; 16 Q; 25 E; 0 Z;
                 28 G; 14 H; 16 I; 48 L; 20 K; 9 M; 17 F; 44 P; 31 S;
                 29 T; 5 W; 8 Y; 36 V;
SEQUENCE LENGTH:
                 461
SEQUENCE
        1 mglpivpgll lslvllallm gihpsgvtgl vpslgdrekr dnlcpqgkya
       51 hpknnsicct kchkgtylvs dcpspgqetv celshkgtft asqnhvrqcl
      101 scktcrkemf qveispckad mdtvcgckkn qfqrylseth fqcvdcspcf
      151 ngtvtipcke kgntvcncha gfflsgnect pcshckknge cmklclppva
      201 nvtnpqdsgt avllplvifl glcllffici sllcrypqwr prvysiicrd
      251 sapvkevege givtkpltpa sipafsanpg fnptlgfstt prfshpvsst
     301 pispvfgpsn whnfvppvre vvptqgadpl lygclnpvpi papvrkwedv
      351 vaaqpqrldt adpamlyavv dgvpptrwke fmrllglseh eierlelqng
      401 rclreahysm leawrrrtpr deatldvvgr vlcdmnlrgc leniretles
      451 pahsstthlp r
ALIGN Smith-Waterman score: 89
      99 aa overlap starting at 7
     pssclllilipllqlinpgstqcsldsvmdkkikdvlnsleyspspiskklscasv_ksq
      : .... :.. :.
     pglllslvllallmgihpsgvtglvpslgdrekrdnlcpqgkyahpknnsicctkchkgt
     grpsscpagmavtgcacgygcgswdvqlettchc_qcsv
```

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

ylvsdcpspgqetvcelshk_gtftasqnhvrqclsckt

REFERENCE: Derwent DGene Search Report

FRANZ-BACON, et al., USSN: 09/099,898

Atty. Docket No.: DX0744K

Mouse C18

Answer Count

 ${\tt MKPTLCFLFILVSLFPLIVPGNAQCSFESLVDQRIKEALSRQEPKTISCTSVTSSGRLASCPAGMVVTGCACGYGCGS} \\ {\tt WDIRNGNTCHCQCSVMDWASARCCRMA}$

30

30 ANSWERS FOUND ABOVE A THRESHOLD OF 65

Similarity Score 80 ||||| 111111111111111111111111111111111111 1111111111111111111111111111111111 1111111111111111111111111111111111 11111111111111111111111111111111111 11111111111111111111111111111111111

10

20

40

```
ANSWER 1 OF 30 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 98P-W44299 Protein
                                            DGENE
                  Human serrate-2 gene expression products - used to
TITLE:
                  regulate stem cell differentiation, useful in treating
                  neoplasms, e.g. leukaemia
                  Itoh A; Sakano S
INVENTOR:
                  (ASAH) ASAHI KASEI KOGYO KK
PATENT ASSIGNEE:
                  WO 9802458 Al 980122
                                                    103 pp
PATENT INFO:
APPLICATION INFO: WO 97-JP2414
                                 970711
                  JP 97-124063
                                 970514
PRIORITY INFO:
                                 960716
                  JP 96-186220
PAT. SEQ. LOC:
                  Claim 3; Page 62-68
DATA ENTRY DATE:
                  19 JUN 1998 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  Japanese
                  98-110528 [10]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 98N-V15181
DESCRIPTION:
                  Human serrate 2
                  Human; serrate 2; regulation; stem cell;
KEYWORD:
                  differentiation; neoplasm; leukaemia; endothelial cell;
                  tumour
                  Homo sapiens
ORGANISM:
ABSTRACT:
      The present sequence represents human serrate 2. The present
      invention also describes a method for the preparation of the
      polypeptides, and antibodies binding to the polypeptide and its
      fragments. The polypeptide and its fragments expressed by the
      serrate-2 gene can be used to inhibit stem (especially blood stem)
      cell differentiation and to inhibit endothelial cell growth. They
      may be incorporated in a cell culture media for culturing
      undifferentiated stem cells. They can also be used for treatment of
      neoplasms such as leukaemia. The antibodies can be used for the
      diagnosis of malignant tumours
AMINO ACID COUNTS:87 A; 76 R; 59 N; 76 D; 0 B; 130C; 31 Q; 70 E; 0 Z;
                  143G; 34 H; 30 I; 72 L; 33 K; 5 M; 35 F; 88 P; 75 S;
                  50 T; 26 W; 30 Y; 62 V;
SEQUENCE LENGTH:
                  1212
SEQUENCE
        l mgyfelqlsa lrnvngells gaccdgdgrt traggcghde cdtyvrvclk
       51 eyqakvtptg pcsyghgatp vlggnsfylp pagaagdrar araraggdqd
      101 pglvvipfqf awprsftliv eawdwdndtt pneellierv shagminped
      151 rwkslhfsgh vahlelqirv rcdenyysat cnkfcrprnd ffghytcdqy
      201 gnkacmdgwm gkeckeavck qgcnllhggc tvpgecrcsy gwqgrfcdec
      251 vpypgcvhgs cvepwqcnce tnwggllcdk dlnycgshhp ctnggtcina
      301 epdgyrctcp dgysgrncek aehactsnpc anggschevp sgfechcpsg
      351 wsgptcaldi decasnpcaa ggtcvdqvdg fecicpeqwv gatcqldane
      401 cegkpclnaf scknliggyy cdcipgwkgi nchinvndcr gqcqhggtck
      451 dlvngyqcvc prgfggrhce lerdkcassp chsgglcedl adgfhchcpq
      501 gfsgplcevd vdlcepspcr ngarcynleg dyycacpddf ggkncsvpre
      551 pcpggacrvi dgcgsdagpg mpgtaasgvc gphgrcvsqp ggnfscicds
      601 gftgtychen iddclgqpcr nggtcidevd afrcfcpsgw egelcdtnpn
      651 dclpdpchsr grcydlvndf ycacddgwkg ktchsrefqc daytcsnggt
      701 cydsgdtfrc acppgwkgst cavaknsscl pnpcvnggtc vgsgasfsci
      751 crdgwegrtc thntndcnpl pcynggicvd gvnwfrceca pgfagpdcri
      801 nidecqsspc aygatcvdei ngyrcscppg ragprcqevi gfgrscwsrg
      851 tpfphgsswv edcnscrcld grrdcskvwc gwkpcllagq pealsaqcpl
      901 gqrclekapg qclrppceaw gecgaeepps tpclprsghl dnncarltlh
```

951 fnrdhvpqgt tvgaicsgir slpatravar drllvllcdr assgasalev

SEARCHED ON 26 OCT 1998

1001 avsfspard1 pdssliggaa haivaaitgr gnsslllavt evkvetvvtg
1051 gsstgllvpv lcgafsvlwl acvvlcvwwt rkrrkerers rlpreesann
1101 qwaplnpirn pierpgghkd vlyqcknftp pprradealp gpaghaavre
1151 deededlgrg eedsleaekf lshkftkdpg rspgrpahwa sgpkvdnrav
1201 rsinearyag ke
ALIGN Smith-Waterman score: 94
105 aa overlap starting at 253
fplivpgn_aqcsfeslvdqrikealsrqepkti_sctsvtssgrlascpag
<pre>ypgcvhgscvepwqcncetnwgg11cdkd1nycgshhpctnggtcinaepdqyrctcpdg mvvtgcacg_ygcgswdirngntchcqcsvmdwasarc</pre>
: : : : : : : : : : : : : : : :
ysgrncekaehactsnpcanggschevpsgfechcp_sgwsgptc

ANSWER 2 OF 30 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD **DGENE** ACCESSION NUMBER: 98P-W44298 Protein Human serrate-2 gene expression products - used to TITLE: regulate stem cell differentiation, useful in treating neoplasms, e.g. leukaemia Itoh A; Sakano S **INVENTOR:** (ASAH) ASAHI KASEI KOGYO KK PATENT ASSIGNEE: PATENT INFO: WO 9802458 Al 980122 103 pp APPLICATION INFO: WO 97-JP2414 970711 PRIORITY INFO: JP 97-124063 970514 JP 96-186220 960716 Claim 2; Page 57-62 PAT. SEQ. LOC: DATA ENTRY DATE: 19 JUN 1998 (first entry) DOCUMENT TYPE: Patent LANGUAGE: Japanese 98-110528 [10] OTHER SOURCE: CROSS REFERENCES: N-PSDB: 98N-V15181 Human serrate 2 protein fragment DESCRIPTION: Human; serrate 2; regulation; stem cell; KEYWORD: differentiation; neoplasm; leukaemia; endothelial cell; tumour ORGANISM: Homo sapiens ABSTRACT: The present sequence represents a human serrate 2 protein fragment. The present invention also describes a method for the preparation of the polypeptides, and antibodies binding to the polypeptide and its fragments. The polypeptide and its fragments expressed by the serrate-2 gene can be used to inhibit stem (especially blood stem) cell differentiation and to inhibit endothelial cell growth. They may be incorporated in a cell culture media for culturing undifferentiated stem cells. They can also be used for treatment of neoplasms such as leukaemia. The antibodies can be used for the diagnosis of malignant tumours AMINO ACID COUNTS:72 A; 58 R; 52 N; 68 D; 0 B; 126C; 29 Q; 54 E; 0 Z; 132G; 30 H; 27 I; 59 L; 24 K; 5 M; 31 F; 73 P; 67 S; 47 T; 21 W; 28 Y; 52 V; SEQUENCE LENGTH: SEQUENCE l mgyfelqlsa lrnvngells gaccdgdgrt traggcghde cdtyvrvclk 51 eyqakvtptg pcsyghgatp vlggnsfylp pagaagdrar araraggdqd 101 pglvvipfqf awprsftliv eawdwdndtt pneellierv shagminped 151 rwkslhfsgh vahlelqirv rcdenyysat cnkfcrprnd ffghytcdqy 201 gnkacmdgwm gkeckeavck qgcnllhggc tvpgecrcsy gwqgrfcdec 251 vpypgcvhgs cvepwqcnce tnwggllcdk dlnycgshhp ctnggtcina 301 epdqyrctcp dgysgrncek aehactsnpc anggschevp sgfechcpsg 351 wsgptcaldi decasnpcaa ggtcvdqvdg fecicpeqwv gatcqldane 401 cegkpclnaf scknliggyy cdcipgwkgi nchinvndcr gqcqhggtck 451 dlvngyqcvc prgfggrhce lerdkcassp chsgglcedl adgfhchcpq 501 gfsgplcevd vdlcepspcr ngarcynleg dyycacpddf ggkncsvpre 551 pcpggacrvi dgcgsdagpg mpgtaasgvc gphgrcvsqp ggnfscicds 601 gftgtychen iddclgqpcr nggtcidevd afrcfcpsgw egelcdtnpn 651 dclpdpchsr grcydlvndf ycacddgwkg ktchsrefqc daytcsnggt 701 cydsgdtfrc acppgwkgst cavaknsscl pnpcvnggtc vgsgasfsci 751 crdgwegrtc thntndcnpl pcynggicvd gvnwfrceca pgfagpdcri 801 nidecqsspc aygatcvdei ngyrcscppg ragprcqevi gfgrscwsrg 851 tpfphgsswv edcnscrcld grrdcskvwc gwkpcllagq pealsaqcpl 901 gqrclekapg qclrppceaw gecgaeepps tpclprsghl dnncarltlh

951 fnrdhvpqgt tvgaicsgir slpatravar drllvllcdr assgasalev

```
ANSWER 3 OF 30 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
                                            DGENE
ACCESSION NUMBER: 96P-W05834 Protein
                  Vertebrate Serrate protein and related DNA - used to
TITLE:
                  treat or prevent malignancies characterised by
                  increased Notch activity
                  Artavanis-Tsakonas S; Gray G E; Henrique D M P;
INVENTOR:
                  Ish-Horowicz D; Lewis J H; Mann R S; Myat A M
                  (IMCR) IMPERIAL CANCER RES TECHNOLOGY
PATENT ASSIGNEE:
      (UYYA)
                  UNIV YALE
                                                    161 pp
PATENT INFO:
                  WO 9627610 A1 960912
APPLICATION INFO: WO 96-US3172
                                 960307
                                 950307
PRIORITY INFO:
                  US 95-400159
PAT. SEO. LOC:
                  Claim 5; Page 104-107
DATA ENTRY DATE:
                  28 JAN 1997 (first entry)
DOCUMENT TYPE:
                  Patent
                  English
LANGUAGE:
                  96-425379 [42]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 96N-W05834
DESCRIPTION:
                  Human Serrate-2 (HJ2)
                  Serrate-2; human jagged-2; HJ2; Notch; cell
KEYWORD:
                  differentiation; cell fate; central nervous system;
                  cancer; tissue repair; therapy; diagnosis; antibody
ORGANISM:
                  Homo sapiens
ABSTRACT:
      Human Serrate-1 (W05833) and human Serrate-2 (W05833) are ligands
      for the zygotic neurogenic locus Notch, and are believed to play a
      major role in determining cell fates (differentiation) in the
      central nervous system. Their amino acid sequences were deduced
      from cDNA clones (see also T40090-91) isolated from human foetal
      brain cDNA libraries. The proteins, antibodies raised to them, and
      encoding nucleic acids can be used in the detection of Serrate
      sequences and in the treatment of disorders of cell fate or
      differentiation, partic. cancer, nervous system disorders and in
      tissue repair or regeneration
AMINO ACID COUNTS:85 A; 87 R; 58 N; 64 D; 0 B; 137C; 34 Q; 56 E; 0 Z;
                  149G; 34 H; 32 I; 72 L; 36 K; 10 M; 39 F; 90 P; 86 S;
                  59 T; 30 W; 28 Y; 71 V;
SEQUENCE LENGTH:
                  1257
SEQUENCE
        1 minpedrwks lhfsghvahl elqirvrcde nyysatcnkf crprndffgh
       51 ytcdqygnka cmdgwmgkec keavckqgcn llhggctvpg ecrcsygwqg
      101 rfcdecvpyp gcvhgscvep wqcncetnwg gllcdkdlny cgshhpctng
      151 gtcinaepdq yrctcpdgys grncekaeha ctsnpcangg schevpsgfe
      201 chcpsgwsgp tcaldideca snpcaaggtc vdqvdgfeci cpeqwvgatc
      251 qldanecegk pclnafsckn liggyycdci pgwkginchi nvndcrgqcq
      301 hggtckdlvn gyqcvcprgf ggrhcelerd kcasspchsg glcedladgf
      351 hchcpqgfsg plcevdvdlc epspcrngar cynlegdyyc acpddfggkn
      401 csvprepcpg gacrvidgcg sdagpgmpgt aasgvcgphg rcvsqpggnf
      451 scicdsgftg tycheniddc lgqpcrnggt cidevdafrc fcpsgwegel
      501 cdtnpndclp dpchsrgrcy dlvndfycac ddgwkgktch srefqcdayt
      551 csnggtcyds gdtfrcacpp gwkgstcava knssclpnpc vnggtcvgsg
      601 asfscicrdg wegrtcthnt ndcnplpcyn ggicvdgvnw frcecapgfa
      651 gpdcrinide cqsspcayga tcvdeingyr cscppgragp rcqevigfgr
      701 scwsrgtpfp hgsswvedcn scrcldgrrd cskvwcgwkp cllagqpea1
      751 saqcplgqrc lekapgqclr ppceawgecg aeeppstpcl prsghldnnc
      801 arltlhfnrd hvpqgttvga icsgirslpa travardrll vllcdrassg
      851 asavevavsf spardlpdss liqgaahaiv aaitqrgnss lllavtevkv
      901 etvvtggsst gllvpvlcga fsvlwlacvv lcvwwtrkrr kerersrlpr
```

SEARCHED ON 26 OCT 1998

```
951 eesannqwap lnpirnpier pgghkdvlyq cknftppprr rcpgrpatrp
1001 sgrmrrtril aavrrtpwrr rssshtnspk ilaarrggrp tgpqapkwtt
1051 arsgasmrpa tsarevgrlq lgrdpgpsvg ampsagpggr ghvhsffilc
1101 kkttknknqm fifyvsltly klfsncqaen ngvfsdscyf ckvavrgtrc
1151 mkgeskgclr rhqivafvtr gcalftessf ysslgflcap gqsagethgc
1201 vgvahgcwwd pwlmvwpvav ggtrgcqwdl wlsvgptvvg gtlvidvala
1251 agtargc
```

FEATURE TABLE:

Key	Location	Qualifier	 +
Domain		note 	Extracellular_domain a deletion in the encoding cDNA clone results in loss of part of the Serrate-2 signal peptide and beginning of the DSL domain
Domain	•	-	DSL "region of homology with Drosophila Delta and Serrate, predicted to mediate binding with Notch"
Domain	•	1abe1 note 	ELR "epidermal growth factor-like repeat domain"
Region	175105	label	ELR1
Region	•	label	ELR2
Region	141180	label	ELR3
Region	1181218		ELR4
Region	1219256	label	ELR5
Region	257294	label	ELR6
Region	1295331	1abel	ELR7
Region	1332369	1abe1	ELR8
Region	370407	1abe1	ELR9
Region	408435	1abel	Partial_ELR
Region	1436469	1abe1	Partial_ELR
Region	470507	1abel	ELR10
Region	508545	1abe1	ELR11
Region	546584	1abel	ELR12
Region	585622	1abel	ELR13
Region	1623660	1abe1	ELR14
Region	664701	label	ELR15
Region	702718	label	Partial_ELR
Region	719735	labe1	Partial_ELR
Domain	913933	1abe1	Transmembrane_domain
Domain	19341257	1abe1	Intracellular_domain

SEARCHED ON 26 OCT 1998

```
L8
      ANSWER 4 OF 30 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 98P-W54234 peptide
                                            DGENE
TITLE:
                  Detection and therapy of cervical cancer - using
                  specific cervical cancer-associated proteins as targets
                  for treatment or as indicators for detection
INVENTOR:
                  Keesee S K; Obar R; Wu Y
PATENT ASSIGNEE:
                  (MATR-N)MATRITECH INC
PATENT INFO:
                  WO 9809170 A2 980305
                                                     79 pp
APPLICATION INFO: WO 97-US14526 970819
PRIORITY INFO:
                  US 96-705660
                                 960830
PAT. SEQ. LOC:
                  Claim 12; Page 55-56
DATA ENTRY DATE:
                  10 AUG 1998 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
OTHER SOURCE:
                  98-230271 [20]
DESCRIPTION:
                  Human TDP-43 protein
KEYWORD:
                  Cervical cancer-associated protein; CvC; tryptic
                  peptide; human; detection; treatment; TDP-43; TAR DNA
                  binding protein; non-chromatin protein
ORGANISM:
                  Homo sapiens
ABSTRACT:
      This protein is the human TDP-43 protein (also known as TAR DNA
      binding protein) which is used to obtain tryptic peptides which are
      used in a method for detecting cervical cancer. The method involves
      detecting the presence of a cervical cancer-associated protein
      (CvC) in a tissue or body fluid sample. The CvC is characterised as
      having a molecular weight of 44900-69400 Daltons as determined by
      sodium dodecyl-sulphate (SDS)-PAGE techniques and an isoelectric
      point (pI) of 5.1-6.6 as determined by standard isoelectric
      focusing techniques. The protein is further characterised as being
      a non-chromatin protein which is detectable at a higher level in a
      human cervical cancer cell than in a normal human cervical cell, as
      determined by 2D-gel electrophoresis. The methods can be used for
      the early and rapid detection of cervical cancer, for treating
      cervical cancers and for monitoring the efficacy of such treatment
AMINO ACID COUNTS: 26 A; 20 R; 28 N; 22 D; 0 B; 6 C; 24 Q; 22 E; 0 Z;
                  55 G; 5 H; 14 I; 21 L; 20 K; 18 M; 22 F; 16 P; 41 S;
                  15 T; 6 W; 8 Y; 25 V;
SEQUENCE LENGTH:
                  414
SEQUENCE
        1 mseyirvted endepieips eddgtvllst vtaqfpgacg lryrnpvsqc
       51 mrgvrlvegi lhapdagwgn lvyvvnypkd nkrkmdetda ssavkvkrav
      101 qktsdlivlg lpwktteqdl keyfstfgev lmvqvkkdlk tghskgfgfv
      151 rfteyetqvk vmsqrhmidg rwcdcklpns kqsqdeplrs rkvfvgrcte
      201 dmtedelref fsqygdvmdv fipkpfrafa fvtfaddqia qslcgedlii
      251 kgisvhisna epkhnsnrql ersgrfggnp ggfgnqggfg nsrgggaglg
      301 nnqgsnmggg mnfgafsinp ammaaaqaal qsswgmmgml asqqnqsgps
      351 gnnqnqgnmq repnqafgsg nnsysgsnsg aaigwgsasn agsgsgfngg
      401 fgssmdskss gwgm
ALIGN Smith-Waterman score: 80
      74 aa overlap starting at 224
      kptlcflfilvslfplivpgnaqcsfeslvdqrikealsrqepktisctsvtssgrlasc
                         ... :. :.:. . :: ::: : .. :::...
           :::::
      kpfrafaf__v_tfaddqiaqslcg_edliikgisvhisnaepkhnsnrqlersgrfggn
      pagmvvtgcacgyg
      :.:.
            :
```

SEARCHED ON 26 OCT 1998

pggf___gnqqqfq

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

```
ANSWER 5 OF 30 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 94P-R62046 protein
                                            DGENE
TITLE:
                  Treating insulin-like growth factor related disease -
                  by admin. of opt. modified binding protein, also new
                  non-phosphorylated binding protein, e.g. for treating
                  cancer or restenosis
INVENTOR:
                  Cox G N; Russell D A
PATENT ASSIGNEE:
                  (SYND) SYNERGEN INC
PATENT INFO:
                  WO 9422466 A 941013
                                                     68 pp
APPLICATION INFO: WO 94-US3755
                                 940406
                  US 93-45265
                                 930407
PRIORITY INFO:
PAT. SEQ. LOC:
                  Disclosure; Page 10
DATA ENTRY DATE:
                  09 MAY 1995 (first entry)
DOCUMENT TYPE:
                  Patent
                  English
LANGUAGE:
OTHER SOURCE:
                  94-332814 [41]
DESCRIPTION:
                  Human mature IGFBP-1
KEYWORD:
                  IGFBP-1; insulin-like growth factor binding protein-1;
                  IGF-binding protein; BP-1; breast cancer; colon cancer;
                  lung cancer; ovary cancer; liver cancer; osteosarcoma;
                  glioma; rhabdomyosarcoma; restenosis; acromegaly;
                  obesity; diabetic nephropathy; retinopathy
ORGANISM:
                  Homo sapiens
ABSTRACT:
      Muteins of human IGFBP-1 (mature protein given in R62046, signal
      peptide in R62047) have been produced containing a S98C or S101C
      substitution. Such muteins can be attached via thiol-reactive
      groups to PEG to form novel therapeutic agents
AMINO ACID COUNTS:26 A; 10 R; 9 N; 7 D; 0 B; 18 C; 9 Q; 22 E; 0 Z;
                  17 G; 6 H; 9 I; 17 L; 9 K; 3 M; 5 F; 17 P; 21 S; 9
                  T; 5 W; 6 Y; 9 V;
SEQUENCE LENGTH:
                  234
SEQUENCE
        l apwqcapcsa eklalcppvs ascsevtrsa gcgccpmcal plgaacgvat
       51 arcarglscr alpgeqqplh altrgqgacv qesdasapha aeagspespe
      101 steiteeell dnfhlmapse edhsilwdai stydgskalh vtnikkwkep
      151 crielyrvve slakaqetsg eeiskfylpn cnkngfyhsr qcetsmdgea
      201 glcwcvyfwn gkripgspei rgdpncqiyf nvqn
ALIGN Smith-Waterman score: 87
      37 aa overlap starting at 11
      ealsrqepktisctsvtssgrlascpagmvvtgcacg
      : :. : . ::. :: . . :: . : :::
```

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

eklalcppvsascsevtrsagcgccpmcalplgaacg

REFERENCE: Derwent DGene Search Report

FRANZ-BACON, et al., USSN: 09/099,898

Atty. Docket No.: DX0744K

40

50

Mouse C19

 ${\tt MKNLSFPLLFLFPLVPELLGSSMPLCPIDEAIDKKIKQDFNSLFPNAIKNIGLNCWTVSSRGKLASCPEGTAVLSCSC}$ GSACGSWDIREEKVCHCQCARIDWTAARCCKLQVAS

29 ANSWERS FOUND ABOVE A THRESHOLD OF

Similarity Score 91 I \mathbf{I} 11111111 11111111111111111 11111111111111111111111111 111111111111111111111111111111111 1111111111111111111111111111111111 111111111111111111111111111111111 30 10 20 Answer Count

```
ANSWER 1 OF 29 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
L10
                                           DGENE
ACCESSION NUMBER: 95P-R74680 protein
                  Genetically engineered tissue plasminogen activator -
TITLE:
                  is modified at positions 44-50 and 296-302 and is
                  non-glycosylated in Kl and K2 regions, has extended
                  half-life and PAI-1 resistance.
                  Huang C; Huang P; Liu S
INVENTOR:
PATENT ASSIGNEE: (BIOE-N) BIOENGINEERING INST ACAD MILITARY
PATENT INFO:
                  CN 1082111 A 940216
APPLICATION INFO: CN 93-109234
                                930806
                                930806
PRIORITY INFO:
                 CN 93-109234
PAT. SEQ. LOC:
                  Claim 2; Fig 1 and Page 5
DATA ENTRY DATE:
                  04 JAN 1996 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  Chinese
                  95-162457 [22]
OTHER SOURCE:
                  t-PA mutein (N117Q, N184Q, delta 296-302, 44-50
DESCRIPTION:
                  replaced by KPIAEK)
                  Tissue plasminogen activator; tPA; thrombolytic agent;
KEYWORD:
                  mutein; deglycosylated kringle domain; PAI-1 resistance
ORGANISM:
                  Synthetic
ABSTRACT:
      The sequences given in R74678-R74689 are examples of preferred
      mutant versions of human tPA. In all the muteins, amino acids 296-
      302 of wild-type tPA (involved in interaction with PAI-1) have been
      deleted and the kringle domains have been deglycosylated by substn.
      of Asn 117 in K1 and Asn184 in K2 by Asp residues. Also, amino
      acids 44-50 of wild-type tPA are replaced by a motif which differs
      between different muteins. The modified tPA proteins have prolonged
      half-life, are resistant to PAI-l and have affinity for fibrin;
      they are useful as thrombolytic agents
AMINO ACID COUNTS:33 A; 33 R; 20 N; 28 D; 0 B; 35 C; 28 Q; 27 E; 0 Z;
                42 G; 14 H; 20 I; 39 L; 21 K; 5 M; 16 F; 28 P; 45 S;
                  25 T; 13 W; 24 Y; 23 V;
SEQUENCE LENGTH:
                  519
SEQUENCE
        l syqvicrdek tqmiyqqhqs wlrpvlrsnr veycwcnsgr aqckpiaekc
       51 seprcfnggt cqqalyfsdf vcqcpegfag kcceidtrat cyedqgisyr
      101 gtwstaesga ectnwqssal aqkpysgrrp dairlglgnh nycrnpdrds
      151 kpwcyvfkag kyssefcstp acsegnsdcy fgqgsayrgt hsltesgasc
      201 lpwnsmilig kvytaqnpsa qalglgkhny crnpdgdakp wchvlknrrl
      251 tweycdvpsc stcglrqysq pqfrikgglf adiashpwqa aifaerflcg
      301 gilisscwil saahcfqerf pphhltvilg rtyrvvpgee eqkfevekyi
      351 vhkefdddty dndiallqlk sdssrcaqes svvrtvclpp adlqlpdwte
      401 celsgygkhe alspfyserl keahvrlyps srctsqhlln rtvtdnmlca
      451 gdtrsggpqa nlhdacqgds ggplvclndg rmtlvgiisw glgcgqkdvp
      501 gvytkvtnyl dwirdnmrp
FEATURE TABLE:
               |Location|Qualifier|
|label
                                  |finger_domain
               |1..49
Domain
```

| amino acids 44-50 of F domain

Thave been replaced by the

|sequence KPIAEK*

|E_domain

SEARCHED ON 26 OCT 1998

Domain

150..86

Inote

|label

```
Inote
                                 | growth factor domain |
Domain
              |87..175 |label
                                 [Kringle_1
                                 |"substn. of Asn117 (corresp.
                       Inote
                                 Ito position 116 in this
                                 (mutein) by Asp destroys an
                                 |N-linked glycosylation site"
              |176..274|label
                                 |Kringle_2
Domain
                                 | substn. of Asn184 (corresp.
                       Inote
                                 |to position 183 in this
                                 (mutein) by Asp destroys an
                                 |N-linked glycosylation site"
              |275..519|label
                                 |P_domain
Domain
                       Inote
                                 | amino acids 296-302 of native
                                 | tPA have been deleted; these
                                 |residues are involved in
                                 |interaction with PAI-1"
Disulfide_bond |6..36
Disulfide_bond |34..43
Disulfide_bond |50..61
Disulfide_bond |55..72
Disulfide_bond |74..83
Disulfide_bond |91..172 |
Disulfide_bond |112..154|
Disulfide_bond |143..167|
Disulfide_bond | 179..260 |
Disulfide_bond |200..242|
Disulfide_bond |231..255|
Disulfide_bond |263..387|
Disulfide_bond |299..315|
Disulfide_bond |307..376|
Disulfide_bond |401..476|
Disulfide_bond |433..449|
  Disulfide_bond |466..494|
ALIGN Smith-Waterman score: 91
      84 aa overlap starting at 8
      deaidkkikqdfns1fpnaiknig1ncwtvssrgklascpegtavlscscgsacgswdir
      dektqmiyqqhqswlrpv1rsnrveycwcnsgraqckpiaekcseprcfnggtc_qqa1y
      eekvchcqcaridwtaarccklqv
           ::: . :..::....
      fsdfv_cqcp__egfagkcceidt
```

L10 ANSWER 2 OF 29 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 97P-W19616 Protein DGENE

TITLE: New active mutants of radish antifungal protein 2 -

used to generate fungus-resistant plants or as

therapeutic or preservative agents

INVENTOR: Brockaert W F; De Samblanx G W; Rees S B

PATENT ASSIGNEE: (ZENE) ZENECA LTD

PATENT INFO: WO 9721814 A1 970619 39 pp

APPLICATION INFO: WO 96-GB3065 961212 PRIORITY INFO: GB 95-25474 951213

PAT. SEQ. LOC: Claim 1; Fig 1

DATA ENTRY DATE: 13 DEC 1997 (first entry)

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 97-332785 [30]

DESCRIPTION: Radish antifungal protein 2 (Rs-AFP2)

KEYWORD: Rs-AFP2; radish antifungal protein 2; fungicide; salt

tolerance; preservative; transgenic plant; crop

protection

ORGANISM: Raphanus sativus

ABSTRACT:

This polypeptide comprises radish antifungal protein 2 (Rs-AFP2). Novel potent antifungal proteins (see W26371-90) based on Rs-AFP2 contain at least 1 mutation selected from a basic residue at positions 9 or 39, and a hydrophobic residue at positions 5 or 16. Proteins containing Gln5Met (see W26379), Gly16Met (W26380), Gly9Arg (W26376), Val39Arg (W26377) or Gly9Arg plus Val39Arg (W26378) mutations are specifically claimed. A cDNA clone encoding Rs-AFP2 preprotein can be modified by recombinant DNA methods to allow expression of mutant isoforms in yeast as mating factor alpha 1 fusion proteins. The Rs-AFP2 mutants have enhanced salt tolerant antifungal activity, especially when expressed in plant tissue where that may have curative as well as protective effects. They are useful for combating fungal diseases in agricultural, pharmaceutical or preservative applications

AMINO ACID COUNTS:3 A; 3 R; 5 N; 0 D; 0 B; 8 C; 3 Q; 1 E; 0 Z; 4 G; 2 H; 2 I; 2 L; 4 K; 0 M; 2 F; 3 P; 3 S; 1 T; 1 W; 2 Y; 2 V;

SEQUENCE LENGTH: 51

SEQUENCE

1 qklcqrpsgt wsgvcgnnna cknqcirlek arhgscnyvf pahkcicyfp 51 c

FEATURE TABLE:

Key	Location	Qualifier	:
Misc_difference	+====== 5 	note l	"Gln at position 5 may be replaced by a hydrophobic amino acid, preferably Met" "
Misc_difference	9 	note 	"Gly at position 9 may be replaced by a basic amino acid, preferably Arg""
Misc_difference	16 	note 	"Gly at position 16 may be replaced by a hydrophobic amino acid, preferably Met""
Misc_difference	139	Inote	*Val at position 39 may be

|replaced by a basic amino |acid, preferably Arg""

SEARCHED ON 26 OCT 1998

```
ANSWER 3 OF 29 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
L10
ACCESSION NUMBER: 97P-W19281 Protein
                                            DGENE
                  Antifungal peptide derived from radish antifungal
TITLE:
                  protein 2 - and related DNA, useful for producing
                  plants with increased fungal resistance and as
                  therapeutic or preservative agent
                  Borremans F A M; Broekaert W F; De Samblanx; Fant F;
INVENTOR:
                  Meloen R H; Puijk W C; Rees S B; Schaaper W M M;
                  Sijtsma L; Van Amerongen A; Van Gelder W M J
PATENT ASSIGNEE:
                  (ZENE) ZENECA LTD
                  WO 9721815 A2 970619
                                                      65 pp
PATENT INFO:
APPLICATION INFO: WO 96-GB3068
                                 961212
PRIORITY INFO:
                  GB 96-6552
                                 960328
```

GB 95-25455 951213 PAT. SEQ. LOC: Disclosure; Figure 1

DATA ENTRY DATE: 21 JAN 1998 (first entry)

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 97-332786 [30]

DESCRIPTION: Raphanus sativus antifungal protein 2 (Rs-AFP2)

KEYWORD: Antifungal protein; candida; fungal resistance; food additive; radish; crop protection; plant defensin;

bacterial protection; preservative

ORGANISM: Raphanus sativus ABSTRACT:

This protein sequence is the Rhapanus sativus (radish) mature antifungal protein (Rs-AFP2), which is homologous to proteins W19280-W19290. Shorter peptides, based on these proteins have been produced (see W19291-92, W19294-98, W19301-304, W19330-34 and W31765-834). Plants containing DNA sequences encoding these proteins have improved resistance to fungi. Compositions containing the peptides can be used to control fungi or bacteria in pharmaceutical (e.g. treatment of Candida infections) or preservative purposes (as food additives). In agriculture, the peptide may be used to improve disease resistance or disease tolerance of crops, either pre or post harvest. When applied to plants they may also have curative as well as protective actions. The peptides may also be used to protect plants by introducing them, or a microorganism capable of expressing the peptide into the soil

AMINO ACID COUNTS:3 A; 3 R; 5 N; 0 D; 0 B; 8 C; 3 Q; 1 E; 0 Z; 4 G; 2 H; 2 I; 2 L; 4 K; 0 M; 2 F; 3 P; 3 S; 1 T; 1 W; 2 Y; 2 V;

SEQUENCE LENGTH: 51

SEQUENCE

1 qklcqrpsgt wsgvcgnnna cknqcirlek arhgscnyvf pahkcicyfp 51 c

ALIGN Smith-Waterman score: 88
47 aa overlap starting at 2

klcqrpsgtwsgvcgnnnacknqcirlekarhgscnyv_fpahkcic

SEARCHED ON 26 OCT 1998 FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

6

L10 ANSWER 4 OF 29 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 97P-W26380 Protein DGENE

TITLE: New active mutants of radish antifungal protein 2 -

used to generate fungus-resistant plants or as

therapeutic or preservative agents

INVENTOR: Broekaert W F; De Samblanx G W; Rees S B

PATENT ASSIGNEE: (ZENE) ZENECA LTD

PATENT INFO: WO 9721814 A1 970619 39 pp

APPLICATION INFO: WO 96-GB3065 961212 PRIORITY INFO: GB 95-25474 951213

PAT. SEQ. LOC: Claim 2; Page 4

DATA ENTRY DATE: 13 DEC 1997 (first entry)

DOCUMENT TYPE: Patent
LANGUAGE: English

OTHER SOURCE: 97-332785 [30]

DESCRIPTION: Radish antifungal protein 2 mutant (G16M)

KEYWORD: Rs-AFP2; radish antifungal protein 2; fungicide; salt

tolerance; preservative; transgenic plant; crop

protection

ORGANISM: Chimeric Raphanus sativus; Chimeric synthetic

ABSTRACT:

This polypeptide comprises a specifically claimed Gly16Met mutant of radish antifungal protein 2 (Rs-AFP2) (see also W19616) that shows enhanced salt tolerant antifungal activity compared to the wild-type protein, especially when expressed in plants. A cDNA clone encoding Rs-AFP2 preprotein can be modified by recombinant DNA methods to allow expression of the mutant isoform in yeast as a mating factor alpha 1 fusion protein. Rs-AFP2 mutants (see also W26371-79 and W26381-90) are useful for combating fungal diseases in agricultural, pharmaceutical and preservative applications. When applied to plants, they may have curative as well as protective effects

AMINO ACID COUNTS:3 A; 3 R; 5 N; 0 D; 0 B; 8 C; 3 Q; 1 E; 0 Z; 3

G; 2 H; 2 I; 2 L; 4 K; 1 M; 2 F; 3 P; 3 S; 1

T; 1 W; 2 Y; 2 V;

SEQUENCE LENGTH: 51

SEQUENCE

1 qklcqrpsgt wsgvcmnnna cknqcirlek arhgscnyvf pahkcicyfp

51 c

ALIGN Smith-Waterman score: 86

47 aa overlap starting at 2

klascpegtavlscscgsacgswdireekvchcqcaridwtaarc_c

klcqrpsgtwsgvcmnnnacknqcirlekarhgscnyv_fpahkcic

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

7

L10 ANSWER 5 OF 29 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 97P-W26379 Protein DGENE

TITLE: New active mutants of radish ant

New active mutants of radish antifungal protein 2 - used to generate fungus-resistant plants or as

therapeutic or preservative agents

INVENTOR: Broekaert W F; De Samblanx G W; Rees S B

PATENT ASSIGNEE: (ZENE) ZENECA LTD

PATENT INFO: WO 9721814 A1 970619 39 pp

APPLICATION INFO: WO 96-GB3065 961212 PRIORITY INFO: GB 95-25474 951213

PAT. SEQ. LOC: Claim 2; Page 4

DATA ENTRY DATE: 13 DEC 1997 (first entry)

DOCUMENT TYPE: Patent

LANGUAGE: English

OTHER SOURCE: 97-332785 [30]

DESCRIPTION: Radish antifungal protein 2 mutant (Q5M)

KEYWORD: Rs-AFP2; radish antifungal protein 2; fungicide; salt

tolerance; preservative; transgenic plant; crop

protection

ORGANISM: Chimeric Raphanus sativus; Chimeric synthetic

ABSTRACT:

This polypeptide comprises a specifically claimed Gln5Met mutant of radish antifungal protein 2 (Rs-AFP2) (see also W19616) that shows enhanced salt tolerant antifungal activity compared to the wild-type protein, especially when expressed in plants. A cDNA clone encoding Rs-AFP2 preprotein can be modified by recombinant DNA methods to allow expression of the mutant isoform in yeast as a mating factor alpha 1 fusion protein. Rs-AFP2 mutants (see also W26371-78 and W26380-90) are useful for combating fungal diseases in agricultural, pharmaceutical and preservative applications. When applied to plants, they may have curative as well as protective effects

AMINO ACID COUNTS:3 A; 3 R; 5 N; 0 D; 0 B; 8 C; 2 Q; 1 E; 0 Z; 4 G; 2 H; 2 I; 2 L; 4 K; 1 M; 2 F; 3 P; 3 S; 1

T; 1 W; 2 Y; 2 V;

SEQUENCE LENGTH: 51

SEQUENCE

1 qklcmrpsgt wsgvcgnnna cknqcirlek arhgscnyvf pahkcicyfp 51 c

ALIGN Smith-Waterman score: 86

47 aa overlap starting at 2

klascpegtavlscscgsacgswdireekvchcqcaridwtaarc_c

klcmrpsgtwsgvcgnnnacknqcirlekarhgscnyv_fpahkcic

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

8

REFERENCE: Derwent DGene Search Report

FRANZ-BACON, et al., USSN: 09/099,898

Atty. Docket No.: DX0744K

Rat C19

 ${\tt MKNLSFLLLFLFLVLGLLGPSMSLCPMDEAISKKINQDFSSLLPAAMKNTVLHCWSVSSRGRLASCPEGTTVTSCSCGSGCGSWDVREDTMCHCQCGSIDWTAARCCTLRVGS$

333 ANSWERS FOUND ABOVE A THRESHOLD OF 68

Similarity Score 135 | 280 350 210 Answer Count 70 140

```
ANSWER 1 OF 333 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 87P-P70641 protein
                                           DGENE
                 New modified tissue plasminogen activator - with new
TTTLE:
                 encoding DNA, new DNA expression vector, useful for
                 treating vascular disorders, eg, pulmonary embolism
                 arterial thrombosis
                 Bang N U; Little S P; Schoner B E; Weigel B J
INVENTOR:
                 (ELIL) ELI LILLY & CO
PATENT ASSIGNEE:
PATENT INFO:
                 AU 8661804 A 870305
                                                   164 pp
APPLICATION INFO: AU 86-61804
                                860825
                 US 85-769298
                                850826
PRIORITY INFO:
                 Claim 9; page 122
PAT. SEQ. LOC:
DATA ENTRY DATE: 10 APR 1991 (first entry)
DOCUMENT TYPE:
                 Patent
                 English
LANGUAGE:
                 87-108842 [16]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 87N-N70990
                 Modified tissue plasminogen activator
DESCRIPTION:
                 Tissue plasminogen activator; kringle domain; embolism;
KEYWORD:
                 thrombosis; stroke;
ORGANISM:
                 Homo sapiens
ABSTRACT:
     The modified t-PA has all/part of the kringle domains of native
     t-PA removed. The t-PA has functional properties superior to those
     of native t-PA. It retains fibrin binding properties and interacts
     more slowly and inefficiently with plasminogen activator
      inhibitor(s) compared to native t-PA. It is obtd. in large amts.
      from a prokaryotic host. Modified t-PA used for treating vascular
      disorders, eg deep vein thrombosis, pulmonary embolism, peripheral
      arterial thrombosis, disseminated intravascular coagulation, emboli
      from the heart or peripheral arteries, acute myocardial infarction,
      thrombotic strokes or fibrin deposits associated with invasive
      cancers. t-PA is used at a dosage of 250000 to 5000000 units at a
      loading dose or in a deep vein thrombosis-pulmonary embolism, or
      250000-7500000 units over 30-90 mins. in acute myocardial
      infarction
AMINO ACID COUNTS:18 A; 24 R; 9 N; 19 D; 0 B; 23 C; 22 Q; 19 E; 0 Z;
                 28 G; 12 H; 15 I; 30 L; 13 K; 5 M; 13 F; 19 P; 29 S;
                 16 T; 7 W; 13 Y; 20 V;
SEQUENCE LENGTH:
                 354
SEQUENCE
       l mgsyqvicrd ektqmiyqqh qswlrpvlrs nrveycwcts graqchsvpv
       51 kscseprcfn ggtcqqalyf sdfvcqcpeg fagkcceist cglrqysqpq
      101 frikgglfad iashpwqaai fakhrrspge rflcggilis scwilsaahc
      151 fgerfpphhl tvilgrtyrv vpgeeeqkfe vekyivhkef dddtydndia
      201 llqlksdssr caqesslvrt vclppadlql pdwtecelsg ygkhealspf
      251 yserlkeahv rlypssrcts qhllnrtvtd nmlcagdtrs ggpqanlhda
      301 cqgdsggplv clndgrmtlv giiswglgcg qkdvpgvytk vtnyldwird
      351 nmrp
ALIGN Smith-Waterman score: 135
      93 aa overlap starting at 1
      lgpsmslcpmdeaiskkinqdfssllpaamkntvlhcwsvssrgrlascp_egttvtscs
      mgsyqvic_rdektqmiyqqhqswlrpvlrsnrveycwctsgraqchsvpvkscseprcf
      cgsgcgswdvredtmchcqcgsidwtaarcct1
       :.:.
                 : .:.: :
```

SEARCHED ON 26 OCT 1998

FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

nggtcqqalyfsdfvcqcpeg___fagkccei

```
ANSWER 2 OF 333 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 95P-R74682 protein
                                            DGENE
                  Genetically engineered tissue plasminogen activator -
TITLE:
                  is modified at positions 44-50 and 296-302 and is
                  non-glycosylated in K1 and K2 regions, has extended
                  half-life and PAI-1 resistance
                  Huang C; Huang P; Liu S
INVENTOR:
                  (BIOE-N) BIOENGINEERING INST ACAD MILITARY
PATENT ASSIGNEE:
PATENT INFO:
                  CN 1082111 A 940216
                                 930806
APPLICATION INFO: CN 93-109234
                  CN 93-109234
                                 930806
PRIORITY INFO:
                  Claim 2; Fig 1 and Page 5
PAT. SEQ. LOC:
                  04 JAN 1996 (first entry)
DATA ENTRY DATE:
                  Patent
DOCUMENT TYPE:
                  Chinese
LANGUAGE:
                  95-162457 [22]
OTHER SOURCE:
                  t-PA mutein (N117Q, N184Q, delta 296-302, 44-50
DESCRIPTION:
                  replaced by ERHTSVQT)
                  Tissue plasminogen activator; tPA; thrombolytic agent;
KEYWORD:
                  mutein; deglycosylated kringle domain; PAI-1 resistance
ORGANISM:
                  Synthetic
ABSTRACT:
      The sequences given in R74678-R74689 are examples of preferred
      mutant versions of human tPA. In all the muteins, amino acids 296-
      302 of wild-type tPA (involved in interaction with PAI-1) have been
      deleted and the kringle domains have been deglycosylated by substn.
      of Asn 117 in K1 and Asn184 in K2 by Asp residues. Also, amino
      acids 44-50 of wild-type tPA are replaced by a motif which differs
      between different muteins. The modified tPA proteins have prolonged
      half-life, are resistant to PAI-1 and have affinity for fibrin;
      they are useful as thrombolytic agents
AMINO ACID COUNTS:32 A; 34 R; 20 N; 28 D; 0 B; 35 C; 29 Q; 27 E; 0 Z;
                  42 G; 15 H; 19 I; 39 L; 19 K; 5 M; 16 F; 27 P; 46 S;
                  27 T; 13 W; 24 Y; 24 V;
SEOUENCE LENGTH:
SEOUENCE
        1 syqvicrdek tqmiyqqhqs wlrpvlrsnr veycwcnsgr aqcerhtsvq
       51 tcseprcfng gtcqqalyfs dfvcqcpegf agkcceidtr atcyedqgis
      101 yrgtwstaes gaectnwgss alagkpysgr rpdairlglg nhnycrnpdr
      151 dskpwcyvfk agkyssefcs tpacsegnsd cyfgqgsayr gthsltesga
      201 sclpwnsmil igkvytaqnp saqalglgkh nycrnpdgda kpwchvlknr
      251 rltweycdvp scstcglrqy sqpqfrikgg lfadiashpw qaaifaerfl
      301 cggilisscw ilsaahcfqe rfpphhltvi lgrtyrvvpg eeeqkfevek
      351 yivhkefddd tydndiallg lksdssrcag essvvrtvcl ppadlglpdw
      401 tecelsgygk healspfyse rlkeahvrly pssrctsqhl lnrtvtdnml
      451 cagdtrsggp qanlhdacqg dsggplvcln dgrmtlvgii swglgcgqkd
      501 vpgvytkvtn yldwirdnmr p
```

FEATURE TABLE:

Key	· · · · · · · · · · · · · · · · · · ·	n Qualifi		
========	:====+======	=+======	==+========	
Domain	151	label	finger_domain	*
	1	note	amino acids 44-50 c	of F domain
	ĺ	ĺ	have been replaced	by the
	İ	ĺ	sequence ERHTSVQT"	
Domain	15288	label	E_domain	

SEARCHED ON 26 OCT 1998

```
| growth factor domain |
                       Inote
Domain
               |89..177 |1abe1
                                 [Kringle_1
                                 | substn. of Asn117 (corresp.
                       Inote
                                 Ito position 118 in this
                                 [mutein] by Asp destroys an
                                 |N-linked glycosylation site"
                                 |Kringle_2
Domain
               |178..276|1abe1
                                 |"substn. of Asn184 (corresp.
                       Inote
                                 |to position 185 in this
                                 (mutein) by Asp destroys an
                                 |N-linked glycosylation site"
               |277..521|1abe1
                                 |P domain
Domain
                                 I amino acids 296-302 of native
                       Inote
                                 |tPA have been deleted; these
                                 Iresidues are involved in
                                 |interaction with PAI-1"
Disulfide_bond | 6...36
Disulfide_bond |34..43
Disulfide_bond |52..63
Disulfide_bond |57..74
Disulfide_bond |76..85
Disulfide_bond |93..174 |
Disulfide_bond |114..156|
Disulfide_bond |145..169|
Disulfide_bond |181..262|
Disulfide_bond |202..244|
Disulfide_bond |233..257|
Disulfide_bond [265..389]
Disulfide_bond |301..317|
Disulfide_bond |309..378|
Disulfide_bond |403..478|
Disulfide_bond |435..451|
   Disulfide_bond |468..496|
ALIGN Smith-Waterman score: 134
      87 aa overlap starting at 8
      deaiskkinqdfssllpaamkntvlhcwsvssrgrlascpegttvtscs___c_gsgcg
      dektqmiyqqhqswlrpvlrsnrveycw_cns_gr_aqcerhtsvqtcseprcfnggtcq
      swdvredtmchcqcgsidwtaarcct1
          : .:.: :
                        :..:: .
      qalyfsdfvcqcpeg___fagkccei
```

```
ANSWER 3 OF 333 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 98P-W54147 protein
                                           DGENE
                 Mutant tissue plasminogen activator proteins - useful
TITLE:
                 for treating vascular disorders, preventing tissue
                 adhesion(s), etc
                 Goeddel D V; Leung D W H; Rice G C
INVENTOR:
                 (GETH) GENENTECH INC
PATENT ASSIGNEE:
                 US 5736135 A 980407
                                                    24 pp
PATENT INFO:
APPLICATION INFO: US 95-389615
                                950213
                 US 91-728456
PRIORITY INFO:
                                910711
                 US 93-8940
                               930126
                 US 94-221660
                                940401
                 US 95-389615
                                950213
                 Claim 6; Page -
PAT. SEQ. LOC:
                 20 JUL 1998 (first entry)
DATA ENTRY DATE:
                 Patent
DOCUMENT TYPE:
LANGUAGE:
                 English
OTHER SOURCE:
                 98-239153 [21]
                 t-PA mutant (Y93C, T103A, N184S, G198D)
DESCRIPTION:
                 Amino acid substitution; t-PA; vascular disorder;
KEYWORD:
                 prevention; fibrin deposition; adhesion formation
ORGANISM:
                 Synthetic
ABSTRACT:
     Mutant tissue plasminogen activator proteins (W54147-W54158) are
      created by single or multiple amino acid substitutions.
      Compositions containing the t-PA variant are used for treating
      vascular disorders, for preventing fibrin deposition or for
      preventing adhesion formation or reformation. Note: This sequence
      is not given in the specification but was created from the wild
      type by the indexer
AMINO ACID COUNTS:33 A; 35 R; 21 N; 29 D; 0 B; 36 C; 26 Q; 26 E; 0 Z;
                  42 G; 16 H; 19 I; 39 L; 21 K; 5 M; 16 F; 29 P; 49 S;
                 24 T; 13 W; 34 Y; 14 V; 0 Others;
SEQUENCE LENGTH:
                 527
SEOUENCE
        1 syqvicrdek tqmiyqqhqs wlrpvlrsnr veycwcnsgr aqchsvpyks
       51 cseprcfngg tcqqalyfsd fvcqcpegfa gkcceidtra tccedqgisy
      101 rgawstaesg aectnwnssa laqkpysgrr pdairlg1gn hnycrnpdrd
      151 skpwcyvfka gkyssefcst pacsegnsdc yfgsgsayrg thsltesdas
      201 clpwnsmili gkvytaqnps aqalglgkhn ycrnpdgdak pwchylknrr
      251 ltweycdyps cstcglrqys qpqfrikggl fadiashpwq aaifakhrrs
      301 pgerflcggi lisscwilsa ahcfqerfpp hhltyilgrt yrvvpgeeeq
      351 kfeyekyiyh kefdddtydn diallqlksd ssrcaqessv vrtvclppad
      401 lqlpdwtece lsgygkheal spfyserlke ahvrlypssr ctsqhllnrt
      451 ytdnmlcagd trsggpqanl hdacqgdsgg plyc1ndgrm tlygiiswgl
      501 gcgqkdypgy ytkvtnyldw irdnmrp
FEATURE TABLE:
               |Location|Qualifier|
______
               11..44
                         Inote
                                  !"Finger domain"
Domain
                                  |"Growth factor domain"
Domain
                |45..91 |note
                                  | "Kringle-1 domain"
Domain
                |92..173 |note
                                  |"Kringle-2 domain"
                |180..261|note
Domain
                                   |"Serine protease domain"
Domain
                |264..527|note
```

|"Y changed from wt to T in

SEARCHED ON 26 OCT 1998

Misc_difference |93

Inote

	1	1	mutant"		
Misc_difference	103 	note 	"T changed mutant"	from wt	to A in
Misc_difference	184 	note 	"N changed mutant"	from wt	to S in
Misc_difference	198	Inote	"G changed	from wt	to D in
	1	1	mutant"	•	•
				-	
ALIGN Smith-Wate	rman scor	e: 129			
87 aa over	lap start	ing at 8	•		
deaiskking	dfss11paa	mkntv1hcws	svssrgrlascp_	_egttvts	cscgsgcgswdv
:::	: : :.	.: : .::	::: : :		: :. : .
dektqmiyqq	hqswlrpvl	rsnrveycwo	cnsgraqchsvpy	ykscsepr	cfnggtcqqaly
redtmchc_	qcgsi	.dwtaarcc			
: .:.:	.: :	: : : ::			
fsdfycacpe	afaakccei	d tratcc			

ANSWER 4 OF 333 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD ACCESSION NUMBER: 95P-R74688 protein **DGENE** TITLE: Genetically engineered tissue plasminogen activator is modified at positions 44-50 and 296-302 and is non-glycosylated in Kl and K2 regions, has extended half-life and PAI-1 resistance **INVENTOR:** Huang C; Huang P; Liu S PATENT ASSIGNEE: (BIOE-N)BIOENGINEERING INST ACAD MILITARY PATENT INFO: CN 1082111 A 940216 APPLICATION INFO: CN 93-109234 930806 PRIORITY INFO: CN 93-109234 930806 Claim 2; Fig 1 and Page 5 PAT. SEQ. LOC: DATA ENTRY DATE: 04 JAN 1996 (first entry) DOCUMENT TYPE: Patent LANGUAGE: Chinese 95-162457 [22] OTHER SOURCE: t-PA mutein (N117Q, N184Q, delta 296-302, 44-50 DESCRIPTION: replaced by DNCRRPG) Tissue plasminogen activator; tPA; thrombolytic agent; KEYWORD: mutein; deglycosylated kringle domain; PAI-l resistance ORGANISM: Synthetic ABSTRACT: The sequences given in R74678-R74689 are examples of preferred mutant versions of human tPA. In all the muteins, amino acids 296-302 of wild-type tPA (involved in interaction with PAI-1) have been deleted and the kringle domains have been deglycosylated by substn. of Asn 117 in K1 and Asn184 in K2 by Asp residues. Also, amino acids 44-50 of wild-type tPA are replaced by a motif which differs between different muteins. The modified tPA proteins have prolonged half-life, are resistant to PAI-1 and have affinity for fibrin; they are useful as thrombolytic agents AMINO ACID COUNTS: 32 A; 35 R; 21 N; 29 D; 0 B; 36 C; 28 Q; 26 E; 0 Z; 43 G; 14 H; 19 I; 39 L; 19 K; 5 M; 16 F; 28 P; 45 S; 25 T; 13 W; 24 Y; 23 V; SEQUENCE LENGTH: 520 SEQUENCE 1 syqvicrdek tqmiyqqhqs wlrpvlrsnr veycwcnsgr aqcdncrrpg 51 cseprcfngg tcqqalyfsd fvcqcpegfa gkcceidtra tcyedqgisy 101 rgtwstaesg aectnwqssa laqkpysgrr pdairlglgn hnycrnpdrd 151 skpwcyvfka gkyssefcst pacsegnsdc yfgqgsayrg thsltesgas 201 clpwnsmili gkvytaqnps aqalglgkhn ycrnpdgdak pwchvlknrr 251 ltweycdvps cstcqlrqys qpqfrikgql fadiashpwq aaifaerflc 301 ggilisscwi lsaahcfger fpphhltvil grtyrvvpge eegkfeveky 351 ivhkefdddt ydndiallql ksdssrcage ssvvrtvclp padlqlpdwt 401 ecelsgygkh ealspfyser lkeahvrlyp ssrctsghll nrtvtdnmlc 451 agdtrsggpq anlhdacqgd sggplvclnd grmtlvgiis wglgcgqkdv 501 pgvytkvtny ldwirdnmrp

FEATURE TABLE:

Key		n Qualifie: =+======		
Domain	150 	label note label	finger_domain "amino acids 44-50 have been replaced sequence DNCRRPG" E_domain	

SEARCHED ON 26 OCT 1998

```
| growth factor domain |
                       note
Domain
              |88..176 |label
                                 [Kringle_1
                                 |"substn. of Asnll7 by Asp
                       |note.
                                 |destroys an N-linked
                                 |glycosylation site"
Domain
             | | 177...275 | label
                                 |Kringle_2
                                 |"substn. of Asn184 by Asp
                       Inote
                                 |destroys an N-linked
                                 |glycosylation site*
                                 |P_domain
Domain
              |276..520|label
                                 | amino acids 296-302 of native
                       note
                                 |tPA have been deleted; these
                                 Iresidues are involved in
                                 |interaction with PAI-1*
Disulfide_bond | 6..36
Disulfide_bond |34..43
Disulfide_bond |51..62
Disulfide_bond | 56..73
Disulfide_bond |75..84
Disulfide_bond |92..173 |
Disulfide_bond |113..155|
Disulfide_bond |144..168|
Disulfide_bond |180..261|
Disulfide_bond |201..243|
Disulfide_bond |232..256| -
Disulfide_bond |264..388|
Disulfide_bond |300..316|
Disulfide_bond |308..377|
Disulfide_bond |402..477|
Disulfide_bond |434..450|
  Disulfide_bond |467..495|
ALIGN Smith-Waterman score: 129
      83 aa overlap starting at 8
      deaiskkinqdfssllpaamkntvlhcwsvssrgrlascpe_gttvtscscgsgcgswdv
      dektqmiyqqhqswlrpvlrsnrveycwcnsgraqcdncrrpgcseprcfnggtcqqaly
     redtmchcqcgsidwtaarcctl
        : .:.: :
                    :..:: .
      fsdfvcqcpeg___fagkccei
```

```
ANSWER 5 OF 333 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 95P-R74678 protein
                                            DGENE
TITLE:
                  Genetically engineered tissue plasminogen activator -
                  is modified at positions 44-50 and 296-302 and is
                  non-glycosylated in Kl and K2 regions, has extended
                  half-life and PAI-1 resistance
INVENTOR:
                  Huang C; Huang P; Liu S
PATENT ASSIGNEE:
                  (BIOE-N) BIOENGINEERING INST ACAD MILITARY
PATENT INFO:
                  CN 1082111 A 940216
APPLICATION INFO: CN 93-109234
                                 930806
PRIORITY INFO:
                  CN 93-109234
                                 930806
PAT. SEQ. LOC:
                  Claim 2; Fig 1 and Page 5
DATA ENTRY DATE:
                  04 JAN 1996 (first entry)
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  Chinese
OTHER SOURCE:
                  95-162457 [22]
DESCRIPTION:
                  t-PA mutein (N117Q, N184Q, delta 296-302, 44-50
                  replaced by ESKPEAEE)
KEYWORD:
                  Tissue plasminogen activator; tPA; thrombolytic agent;
                  mutein; deglycosylated kringle domain; PAI-1 resistance
ORGANISM:
                  Synthetic
ABSTRACT:
      The sequences given in R74678-R74689 are examples of preferred
      mutant versions of human tPA. In all the muteins, amino acids 296-
      302 of wild-type tPA (involved in interaction with PAI-1) have been
      deleted and the kringle domains have been deglycosylated by substn.
      of Asn 117 in K1 and Asn184 in K2 by Asp residues. Also, amino
      acids 44-50 of wild-type tPA are replaced by a motif which differs
      between different muteins. The modified tPA proteins have prolonged
      half-life, are resistant to PAI-1 and have affinity for fibrin;
      they are useful as thrombolytic agents
AMINO ACID COUNTS:33 A; 33 R; 20 N; 28 D; 0 B; 35 C; 28 Q; 30 E; 0 Z;
                  42 G; 14 H; 19 I; 39 L; 20 K; 5 M; 16 F; 28 P; 46 S;
                  25 T; 13 W; 24 Y; 23 V;
SEQUENCE LENGTH:
                  521
SEQUENCE
        1 syqvicrdek tqmiyqqhqs wlrpvlrsnr veycwcnsgr aqceskpeae
       51 ecseprcfng gtcqqalyfs dfvcqcpegf agkcceidtr atcyedqgis
      101 yrgtwstaes gaectnwqss alaqkpysgr rpdair1g1g nhnycrnpdr
      151 dskpwcyvfk agkyssefcs tpacsegnsd cyfgqgsayr gths1tesga
      201 sclpwnsmil igkvytagnp sagalglgkh nycrnpdgda kpwchvlknr
      251 rltweycdvp scstcglrqy sqpqfrikgg lfadiashpw qaaifaerfl
      301 cggilisscw ilsaahcfqe rfpphhltvi 1grtyrvvpg eeeqkfevek
      351 yivhkefddd tydndiallg lksdssrcag essvvrtvcl ppadlglpdw
      401 tecelsgygk healspfyse rlkeahvrly pssrctsghl lnrtvtdnml
      451 cagdtrsggp qanlhdacqq dsqqplvcln dqrmtlvqii swqlqcqqkd
      501 vpgvytkvtn y1dwirdnmr p
FEATURE TABLE:
```

TERIORE IRDEE,

SEARCHED ON 26 OCT 1998

Domain 277521 label P_domain amino acids 296-302 of native tPA have been deleted; these tesidues are involved in linteraction with PAI-1" Disulfide_bond 636 Disulfide_bond 5263 Disulfide_bond 5774 Disulfide_bond 7685 Disulfide_bond 93174 Disulfide_bond 114156 Disulfide_bond 145169 Disulfide_bond 181262 Disulfide_bond 202244 Disulfide_bond 233257
Disulfide_bond 636
Disulfide_bond 3443
Disulfide_bond 5263
Disulfide_bond 5774
Disulfide_bond 7685
Disulfide_bond 93174
Disulfide_bond 114156 Disulfide_bond 145169 Disulfide_bond 181262 Disulfide_bond 202244 Disulfide_bond 233257
Disulfide_bond 145169 Disulfide_bond 181262 Disulfide_bond 202244 Disulfide_bond 233257
Disulfide_bond 181262 Disulfide_bond 202244 Disulfide_bond 233257
Disulfide_bond 202244
Disulfide_bond 233257
. =
D15411146_D0114 120311505
Disulfide_bond 301317
Disulfide_bond 309378
Disulfide_bond 403478
Disulfide_bond 435451
Disulfide_bond 468496
ALIGN Smith-Waterman score: 129
84 aa overlap starting at 8
deaiskkinqdfssllpaamkntvlhcwsvssrgrlascpegttvtscscgsgcgswd
11 1 1 1 1 1 1 . 1 . 1 . 1 .
dektqmiyqqhqswlrpvlrsnrveycwcnsgraqceskpeaeecseprcfnggtcqqal
vredtmchcqcgsidwtaarcctl
1 .1.1 :
yfsdfvcqcpegfagkccei

REFERENCE: Derwent DGene Search Report FRANZ-BACON, et al., USSN: 09/099,898

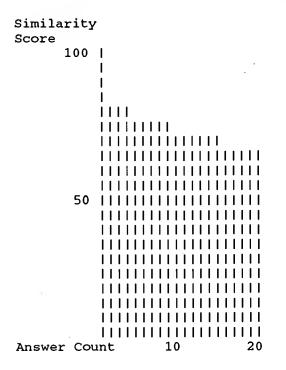
Atty. Docket No.: DX0744K

Human C23

 ${\tt MKALCLLLLPVLGLLVSSKTLCSMEEAINERIQEVAGSLIFRAISSIGLECQSVTSRGDLATCPRGFAVTGCTCGSAC}$ GSWDVRAETTCHCQCAGMDWTGARCCRVQP

30

20 ANSWERS FOUND ABOVE A THRESHOLD OF 66



40

50

```
ANSWER 1 OF 20 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
L14
ACCESSION NUMBER: 97P-W24566 peptide
                                           DGENE
                  Serine protease from Streptomyces griseus ATCC 55178 -
TITLE:
                  with good stability in presence of urea or guanidine,
                  useful in cleaning compositions, including laundry and
                  dishwashing detergents
INVENTOR:
                  Leigh S D
                  (CLRX) CLOROX CO
PATENT ASSIGNEE:
                  US 5646028 A 970708
                                                     16 pp
PATENT INFO:
APPLICATION INFO: US 91-718303
                                 910618
                                 910618
PRIORITY INFO:
                  US 91-718303
                  US 92-973343
                                 921106
                                 940818
                  US 94-292924
                  US 95-544143
                                 951017
                  Claim 3; Column 25
PAT. SEQ. LOC:
                  05 NOV 1997 (first entry)
DATA ENTRY DATE:
DOCUMENT TYPE:
                  Patent
LANGUAGE:
                  English
                  97-362936 [33]
OTHER SOURCE:
                  Serine protease C-terminal sequence
DESCRIPTION:
                  Serine protease; C-terminus; Streptomyces griseus;
KEYWORD:
                  guanidine; pre-soak; cleaning composition; laundry
                  detergent; additive composition; enzyme; dishwasher
                  detergent; drain opener; urea; contact lens cleanser;
                  proteinaceous stain
                  Streptomyces griseus variety alkaliphilus No. 33
ORGANISM:
ABSTRACT:
      This sequence represents the C-terminal sequence of the serine
      protease of the invention. The serine protease was isolated from
      Streptomyces griseus variety alkaliphilus No. 33 (ATCC 55178). The
      protease has an apparent molecular weight of 19 kD (by reducing
      sodium dodecylsulphate polyacrylamide gel electrophoresis), and
      improved stability against urea and guanidine. The serine protease
      is specific for the substrate represented by W24567, but also
      recognises the substrates shown in W26078-W26096. The protease is
      inhibited by phenylmethylsulphonyl fluoride. The serine protease is
      useful in liquid or granular cleaning compositions, specifically
      laundry detergents or additive compositions. It is also useful in
      automatic dishwasher detergents, pre-soaks, drain openers, contact
      lens cleansers etc. The protease has better activity against
      proteinaceous stains than known enzymes and unusually high
      stability in the presence of chaotropic agents
                                                        Q; 0 E; 0 Z;
AMINO ACID COUNTS: 4 A; 4 R; 3 N; 1 D; 0 B; 4 C; 9
                  18 G; 2 H; 6 I; 3 L; 0 K; 0 M; 2 F; 4 P; 15 S;
                  17 T; 1 W; 2 Y; 7 V;
SEQUENCE LENGTH:
                  102
SEQUENCE
        1 vtgstqatvg ssicrsgstt gwrcgtiqqh ntsvtypqgt itgvtrtsac
       51 aqpgdsggsf isgtqaqgvt sggsgncsig gttfhqpvnp ilsqygltlv
      101 rs
   ALIGN Smith-Waterman score: 101
      75 aa overlap starting at 6
      qevagslifraissiglecqsvtsrgdlatcprgfavtgctcgsacgs_wdvraettchc
       : ..:: : :. :. : .: .. .. .: :.: : ::: : ::: :
      qatvgssicrsgsttgwrcgtiqqhntsvtypqg_titgvtrtsacaqpgdsggsfisgt
      qcagmdwtgarccrv
         :. :. : .
      qaqgvtsggsgncsi
```

SEARCHED ON 26 OCT 1998

L14 ANSWER 2 OF 20 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 95P-R77256 Protein DGENE

TITLE: Pure, truncated fungal cellulase protein from

Trichoderma - useful to reduce or eliminate dye, colourant or pigment back-staining or redeposition in

stone-washing or bio-polishing

INVENTOR: Clarkson K A; Collier K D; Fowler T; Larenas E; Ward M

PATENT ASSIGNEE: (GEMV)GENENCOR INT INC

PATENT INFO: WO 9516782 A 950622 105 pp

APPLICATION INFO: WO 94-US14163 941219 PRIORITY INFO: US 93-169948 931217 PAT. SEQ. LOC: Claim 12; Page 38

DATA ENTRY DATE: 13 DEC 1995 (first entry)

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 95-231574 [30] CROSS REFERENCES: N-PSDB: 95N-Q91276

DESCRIPTION: Truncated endoglucanase EGI catalytic core

KEYWORD: Cellulase; catalytic core; enzyme

ORGANISM: Trichoderma longibrachiatum

ABSTRACT:

A truncated fungal cellulase of Trichoderma comprising a EGI catalytic core with the sequence in R77256, which is encoded by Q91276, is claimed. The truncated cellulase is capable of endoglucanase activity. Genes for EGI and EGII have been isolated from T. longibrachiatum and the protein domain structure has been confirmed (Penttila, M. et al. 1986, Gene 45, 253-263; Van Arsdell, J.N. et al. 1987, Bio/Technology 5, 60-64; Saloheimo, M. et al., 1988, Gene 63, 11-21)

AMINO ACID COUNTS:2 A; 0 R; 2 N; 1 D; 0 B; 6 C; 4 Q; 0 E; 0 Z; 5 G; 0 H; 0 I; 1 L; 0 K; 0 M; 0 F; 2 P; 7 S; 2 T; 2 W; 3 Y; 2 V;

SEQUENCE LENGTH: 39

SEQUENCE

1 qacssvwgqc ggqnwsgptc casgstcvys ndyysqc1p ALIGN Smith-Waterman score: 79

13 aa overlap starting at 9

qcagmdwtgarcc
::::::::
qcggqnwsgptcc

SEARCHED ON 26 OCT 1998 FILE LAST UPDATED: 18 OCT 1998 <19981018/UP>

3 :

```
ANSWER 3 OF 20 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 96P-R98208 Protein
                                            DGENE
                  Cell-targetted retroviral vector particles - having
TITLE:
                  envelope protein modified with targetting polypeptide
                  Anderson W; Chiang Y L; Januszeski M; Mackrell A J;
INVENTOR:
                  Zhao Y
                  (GENE-N) GENETIC THERAPY INC
PATENT ASSIGNEE:
                  UNIV SOUTHERN CALIFORNIA
      (UYSC-N)
                                                     73 pp
PATENT INFO:
                  WO 9630504 A1 961003
                                 960322
APPLICATION INFO: WO 96-US3908
                 US 95-409648
                                 950324
PRIORITY INFO:
                  Example 2; Page 36
PAT. SEQ. LOC:
DATA ENTRY DATE:
                 30 DEC 1996 (first entry)
DOCUMENT TYPE:
                  Patent
                  English
LANGUAGE:
OTHER SOURCE:
                  96-455352 [45]
                  Nucleotide used in production of MSH/MoMuLV chimeric
DESCRIPTION:
                  sequence
                  Moloney murine leukaemia virus; gp70; 4070A retrovirus;
KEYWORD:
                  retrovirus; 10A1 murine leukaemia virus; NZB-9-1 murine
                  leukaemia virus; polytropic MX27 provirus; targetted
                  drug delivery; gene therapy; single chain antibody;
                  envelope protein; ss
                  Synthetic
ORGANISM:
ABSTRACT:
      Cell targetted retroviral vector particles can be used in gene
      therapy to deliver a heterologous gene to a target cell for
      expression of a heterologous polypeptide in that cell. The cell
      targetted retroviral vector particles comprise an envelope protein
      which is modified to contain a targetting polypeptide (a single
      chain antibody), or in the case of moloney murine leukaemia virus
      (MoMuLV), alpha melanotropin-stimulating hormone (MSH). Two
      oligonucleotides (R98207, R98208) were used to substitute sequences
      in MoMuLV for MSH sequences. This oligonucleotide was used to
      replace residues G80-P88 of MoMuLV envelope protein (See W04248)
AMINO ACID COUNTS:8 A; 0 R; 0 N; 0 D; 0 B; 17 C; 0 Q; 0 E; 0 Z; 8
                  G; 0 H; 0 I; 0 L; 0 K; 0 M; 0 F; 0 P; 0 S; 11
                  T; 0 W; 0 Y; 0 V;
```

SEQUENCE LENGTH: 44

SEQUENCE

1 catttccgat ggtgcaagcc ggtattaacc tccctcaccc ctcg

ALIGN Smith-Waterman score: 83
43 aa overlap starting at 5

tcprgfavtgctcgsacgswdvraettchcqcagmdwtgarcc

:: . . ::: . : . . : :: : ::

tccgatggtgcaagccggtattaacctccctca____cc

SEARCHED ON 26 OCT 1998

```
ANSWER 4 OF 20 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD
ACCESSION NUMBER: 94P-R45359 Protein
                                           DGENE
                 Transgenic plant contg. cDNA encoding wheat or barley
TITLE:
                 lectin - has insecticidal properties in its leaves
INVENTOR:
                 Raikhel N V
PATENT ASSIGNEE:
                  (UNMS)UNIV MICHIGAN STATE
                                                    26 pp
PATENT INFO:
                 US 5276269 A 940104
APPLICATION INFO: US 92-917665
                                920720
PRIORITY INFO:
                 US 89-406318
                                890912
                 US 92-917665
                                920720
PAT. SEQ. LOC:
                 Disclosure; Fig 6
DATA ENTRY DATE: 06 JUL 1994 (first entry)
DOCUMENT TYPE:
                 Patent
LANGUAGE:
                 English
                 94-016167 [02]
OTHER SOURCE:
CROSS REFERENCES: N-PSDB: 94N-Q54433
                 Wheat germ agglutinin isolectin WGA-D
DESCRIPTION:
KEYWORD:
                 Transgenic plant; leaf; leaves; insecticidal;
                 fungicidal; properties; tobacco; gramineae
ORGANISM:
                 Triticum aestivum L
ABSTRACT:
      The sequence is that of wheat germ agglutinin isolectin A, WGA-D,
      which may be expressed in transgenic plants to provide plants
      (pref. tobacco) having insecticidal and fungicidal properties in
      their leaves
AMINO ACID COUNTS:23 A; 5 R; 11 N; 5 D; 0 B; 32 C; 12 Q; 5 E; 0 Z;
                  42 G; 2 H; 3 I; 9 L; 8 K; 6 M; 6 F; 6 P; 16 S; 9
                 T; 3 W; 7 Y; 3 V;
SEQUENCE LENGTH:
                 213
SEQUENCE
        1 mrkmmstmal tlgaavflaf aaataqaqrc geqgsnmecp nnlccsqygy
       51 cgmggdycgk gcqngacwts krcgsqagga tcpnnhccsq yghcgfgaey
      101 cgagcqggpc radikcgsqs ggklcpnnlc csqwgfcglg sefcgggcqs
      151 gacstdkpcg kdaggrvctn nyccskwgsc gigpgycgag cqsggcdavf
      201 agaitanstl lae
ALIGN Smith-Waterman score:
      76 aa overlap starting at 13
      gslifraissiglecqsvtsrgdlatcp____rgf_avtgctcgsacgswdvraett
                                        :. .. : ::..: .
                  . :
                         .:. ::
      gaavflafaaataqaqrcgeqgsnmecpnnlccsqygycgmggdycgkgcqngacwtskr
      chcqcaqmdwtqarcc
      : : .:
      cgsqaggatcpnnhcc
```

L14 ANSWER 5 OF 20 DGENE COPYRIGHT 1998 DERWENT INFORMATION LTD

ACCESSION NUMBER: 96P-W02025 Protein DGENE

TITLE: Treatment of cellulose-contg. fabrics such as denim,

e.g. stone:washing - using truncated cellulase enzyme to increase abrasion and give reduced redeposition of

dye

INVENTOR: Clarkson K A; Collier K D; Fowler T; Larenas E; Ward M

PATENT ASSIGNEE: (GEMV) GENENCOR INT INC

PATENT INFO: WO 9623928 A1 960808 124 pp

APPLICATION INFO: WO 96-US977 960129
PRIORITY INFO: US 95-382452 950201
PAT. SEQ. LOC: Disclosure; Fig 2A-2E
DATA ENTRY DATE: 28 OCT 1996 (first entry)

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: 96-371466 [37] CROSS REFERENCES: N-PSDB: 96N-T32221

DESCRIPTION: Trichoderma cellobiohydrolase II

KEYWORD: Cellobiohydrolase II; CBHII; cellulase; cellulose;

denim; stonewashing; dye redeposition; backstaining

ORGANISM: Trichoderma longibrachiatum

ABSTRACT:

The amino acid sequences for Trichoderma longibrachiatum cellobiohydrolase I (CBHI) (W02022), CBHII (W02025), endoglucanase I (EGI) (W02029), EGII (W02032) and EGIII (W02034) were deduced from the respective genomic DNA sequences (T32220-24). The CBHI, CBHI, EGI and EGII enzymes have catalytic core domains useful for reducing dye redeposition (backstaining) on cellulose-contg. fabrics such as denim, whilst maintaining or increasing abrasion during stonewashing. Truncated enzymes comprising these catalytic core domains can be obtd. by proteolysis of the complete enzyme or by inserting the appropriate DNA fragment into a vector, using this to transform a Trichoderma sp. host cell, and recovering the recombinant core domain

AMINO ACID COUNTS:60 A; 14 R; 30 N; 21 D; 0 B; 12 C; 21 Q; 10 E; 0 Z; 40 G; 4 H; 18 I; 38 L; 10 K; 5 M; 12 F; 32 P; 47 S;

38 T; 12 W; 20 Y; 27 V;

SEQUENCE LENGTH: 471

SEQUENCE

1 mivgilttla tlatlaasvp leerqacssl wgqcggqnws gptccasgst

51 cvysndyysq clpgaassss straasttsr vspttsrsss atpppgsttt

101 rvppvgsgta tysgnpfvgv tpwanayyas evsslaipsl tgamataaaa

151 vakvpsfmwl dtldktplme qtladirtan knggnyagqf vvid1pdrdc

201 aalasngeys iadggvakyk nyidtirqiv veysdirtl1 viepdslan1

251 vtn1gtpkca naqsayleci nyavtqln1p nvamyldagh agwlgwpanq

301 dpaaq1fanv yknasspral rglatnvany ngwnitspps ytqqnavyne

351 klyihaigpl lanhgwsnaf fitdqgrsgk qptgqqwgd wcnvigtgfg

401 irpsantgds 11dsfvwvkp ggecdgtsds saprfdshca lpdalqpapq

451 agawfqayfv q11tnanpsf 1

FEATURE TABLE:

Key |Location|Qualifier|

Peptide |1..24 |1abel |Sig_peptide | Protein |25..471 |1abel |Mat_protein |

Domain |25..63 |label |Cellulose_binding_domain

SEARCHED ON 26 OCT 1998

gilttlatlatlaasvpleerqacsslwgqcggqnwsgptcc

SEARCHED ON 26 OCT 1998